KEY MESSAGES – ZIKA VIRUS DISEASE

<u>Purpose:</u> This document is for internal and external use. The document contains cleared key messages for use in developing other materials.

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Updated information is in blue.

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BACKGROUND ON ZIKA

- Zika virus was first discovered in a monkey in the Zika Forest of Uganda in 1947.
- Before 2007, at least 14 cases of Zika had been documented, although other cases were likely to have occurred and were not reported.
- Zika outbreaks have probably occurred in many locations. Because the symptoms of Zika are similar to those of many other diseases, many cases may not have been recognized.

OUTBREAK SUMMARY

- Before 2015, Zika virus disease (Zika) outbreaks occurred in areas of Africa, Southeast Asia, and the Pacific Islands.
- On May 7, 2015, the Pan American Health Organization (PAHO) issued an <u>alert</u> regarding the first confirmed Zika virus infections in Brazil.
- Since May 2015, CDC has been responding to increased reports of Zika and has assisted in investigations with PAHO and the Brazil Ministry of Health. The first regional travel notices for Zika in South America and Mexico were posted in December 2015.
- Currently, outbreaks are occurring in many countries and territories.
- On January 22, 2016, CDC activated its <u>Emergency Operations Center</u> (EOC) to respond to outbreaks of Zika
 occurring in the Americas and increased reports of birth defects and Guillain-Barré syndrome in areas affected by
 Zika. On February 8, 2016, CDC elevated its EOC activation to a Level 1, the highest level.
- On February 1, 2016, the World Health Organization (WHO) declared a <u>Public Health Emergency of International</u>
 <u>Concern</u> (PHEIC) because of clusters of microcephaly and other neurological disorders in some areas affected by Zika.
- On February 8, 2016, President Obama announced a request for \$1.8 billion in emergency funds for several agencies to accelerate research into a vaccine and educate populations at risk for disease.

ZIKA & THE UNITED STATES

- Zika is a nationally notifiable disease. State and territorial health departments are encouraged to report
 laboratory-confirmed cases to CDC through ArboNET, the national surveillance system for arboviral
 diseases. Healthcare providers should report cases to their local, state or territorial health department according
 to the laws or regulations for reportable diseases in their jurisdiction.
- Pregnant women with any laboratory evidence of possible Zika virus infection should be reported to the <u>US Zika Pregnancy Registry</u>.
- CDC watches for and reports the number of Zika cases and the areas where Zika is spreading, which will help improve our understanding of how and where Zika is spreading.
- For the most recent case counts, visit CDC's <u>Cases in the United States webpage</u>.

LOCAL TRANSMISSION

- Local transmission means that mosquitoes in the area have been infected with Zika virus and can spread it to people.
- In December 2015, the Commonwealth of Puerto Rico, a United States territory, reported its first confirmed locally transmitted Zika virus case.
 - Puerto Rico continues to report the highest number of Zika virus infections in the US, including the highest number of infections among pregnant women.
- Cases of local transmission have also been confirmed in two other US territories, the United States Virgin Islands, and American Samoa.
- Local transmission of Zika virus has been reported in the United States.

 The Florida Department of Health has identified two areas in Miami-Dade County where Zika is being spread by mosquitoes.

- o CDC is working closely with Florida public health officials to investigate these cases.
- CDC and Florida are issuing travel, testing, and other recommendations for people who traveled to or live in the Florida-designated areas.
- At Florida's request, CDC also sent a CDC Emergency Response Team (CERT) with experts in Zika virus, pregnancy and birth defects, vector control, laboratory science, and risk communication to assist in the response.
- See Advice for People Living in or Traveling to South Florida.
- CDC is not able to predict how much Zika virus will spread in the continental United States.
 - Many areas in the United States have the type of <u>mosquitoes</u> that can become infected with and spread
 Zika virus. Recent outbreaks in the continental United States of chikungunya and dengue, which are
 spread by the same type of mosquito, have been relatively small and limited to a small area.
 - We will maintain and improve our ability to identify and test for Zika and other mosquito-borne diseases.
- Aedes aegypti or Ae. albopictus mosquitoes can cause an outbreak of Zika, if the following happens:
 - o People get infected with the virus.
 - An Aedes mosquito bites the infected person during the period of time when the virus can be found in the
 person's blood, typically only through the first week of infection.
 - The infected mosquito lives long enough for the virus to multiply and for the mosquito to bite another person.
 - The cycle continues multiple times to start an outbreak.

TRAVEL-ASSOCIATED CASES

- A travel-associated (or imported) case means that a person with Zika became infected during travel to an area with active Zika transmission. This includes a traveler becoming infected and anyone infected by that traveler.
 - A person with Zika virus can pass it to his or her sex partners.
 - A case of sexually transmitted Zika in the United States is defined as a person who becomes infected with Zika from having sex with an infected person.
 - Zika virus can be passed from a pregnant woman to her fetus during pregnancy or around the time of delivery. It is not known if Zika virus can be passed to the fetus around the time of conception.
- Most travel-associated cases of Zika in the United States have been in travelers coming from the Caribbean, Puerto Rico, and Central and South America.

ZIKA & PUERTO RICO

- The first local transmission of Zika virus in Puerto Rico was reported on December 31, 2015.
- At the request of Governor Alejandro García Padilla, the US Health and Human Services Secretary Sylvia M. Burwell declared a public health emergency (PHE) for Puerto Rico on August 12, 2016.
 - Through the public health emergency declaration, the government of Puerto Rico can
 - Apply for funding to hire and train unemployed workers to assist in vector control and outreach
 and education efforts through the U.S. Department of Labor's National Dislocated Worker Grant
 program; and
 - Request the temporary reassignment of local public health department or agency personnel who
 are funded through Public Health Service Act programs in Puerto Rico to assist in the Zika
 response.
- Since local transmission of Zika virus was reported in Puerto Rico in late 2015, it has spread to all municipalities.
 - The San Juan metropolitan area has the largest number of residents who have tested positive. Ponce,
 Bayamón, and Caguas metropolitan areas are also highly affected.

• In the coming months, it's possible that more pregnant women in Puerto Rico will become infected with Zika. A recent analysis from CDC estimates that 5,900-10,300 pregnant women might be infected during the Zika virus outbreak in Puerto Rico in 2016.

- Because the full spectrum of health effects that might occur from Zika virus infection during pregnancy is unknown, this range reflects the number of infants at risk for health problems caused by Zika.
- CDC scientists estimated that 100-270 of these infections in pregnant women might lead to microcephaly in infants between mid-2016 and mid-2017.
- In February 2016, the Puerto Rico Department of Health reported the first case of Guillain-Barré syndrome (GBS) in a patient with evidence of recent infection Zika virus.
 - Monthly counts of reported cases of GBS with evidence of Zika virus infection appear to be increasing in Puerto Rico since April 2016.
- Outbreaks of mosquito-borne viruses tend to peak in the late summer and fall in Puerto Rico—in hotter months
 with higher rainfall—raising concern that Zika will continue to spread and increase in the coming months.
- The situation in Puerto Rico warrants urgent, comprehensive action to protect pregnant women.
 - The government of Puerto Rico and its municipalities, and everyone in the community, can implement an integrated mosquito management program that includes reducing places where mosquitoes lay eggs, keeping mosquitoes out of houses, and reducing the populations of adult mosquitoes and mosquito larvae by treating areas with Environmental Protection Agency (EPA)-approved products.
- All pregnant women in Puerto Rico should be assessed for possible Zika virus exposure during each prenatal care visit.
- CDC is working with Puerto Rico to provide services to protect pregnant women from Zika, support pregnant women who have been infected with Zika, increase lab testing, improve mosquito control, and provide access to contraception for women who choose to delay or avoid pregnancy.

SYMPTOMS

- Many people infected with Zika virus won't have symptoms or will only have mild symptoms.
- The sickness is usually mild with symptoms lasting for several days to a week.
- The most common symptoms of Zika virus disease are
 - o Fever
 - o Rash
 - Joint pain
 - Conjunctivitis (red eyes)
- Other symptoms include
 - Muscle pain
 - o Headache
- People usually don't get sick enough to go to the hospital, and they very rarely die of Zika.

DIAGNOSIS

- To diagnose Zika, your doctor or other healthcare provider will ask you about any recent travel and any signs and symptoms you may have. A blood or urine test can confirm a Zika infection.
- Pregnant women who have recently traveled to an area with Zika or whose sex partner lives in or has traveled to
 an area with Zika, should talk to a doctor or other healthcare provider about their risk of Zika virus infection even if
 they don't feel sick.
- Pregnant women should see a doctor or other healthcare provider if they develop a fever, rash, joint pain, or
 conjunctivitis (red eyes) during their trip or within 2 weeks after traveling to an area with Zika. They should tell the
 doctor or other healthcare provider where they traveled. CDC has developed <u>interim guidelines</u> for pregnant

women for Zika. The doctor or other healthcare provider may order specialized blood tests to look for Zika or other similar viral diseases like dengue or chikungunya.

- People who are not pregnant should see a doctor or other healthcare provider if they develop symptoms (fever, rash, joint pain, or red eyes) and live in or have recently traveled to an area with Zika. They should tell the healthcare provider that they traveled to an area with Zika.
- A doctor or other healthcare provider may order blood or urine tests to look for Zika or other similar viral diseases like dengue or chikungunya.
- See <u>Laboratory Testing</u> section for more information on Zika testing.

TRANSMISSION

- Zika virus is spread to people primarily through the bite of an infected Aedes species mosquito (Ae. aegypti and Ae. albopictus).
 - o Mosquitoes that spread Zika virus bite during the day and night.
- A pregnant woman can pass Zika virus to her fetus during pregnancy or around the time of birth. We are studying how Zika affects pregnancies. Zika virus is known to cause microcephaly and other brain anomalies.
- A person with Zika virus can pass it to his or her <u>sex partners</u>.
- There is a strong possibility Zika can be spread through blood transfusion.
- One case of Zika has been confirmed in a person in Utah with no known risk factors; however, the person did
 provide care to another person who had uniquely high amounts of Zika virus in his blood. Although the route of
 transmission is not certain, family contacts should be aware that blood and body fluids of severely ill patients
 might be infectious.
- Transmission of Zika virus infection through <u>breastfeeding</u> has not been documented.
- There is no evidence that Zika is spread through touching, coughing, or sneezing.

VECTOR INFORMATION

- A mosquito can spread Zika virus through bites. Not all people who are infected will get sick.
- There are many species of *Aedes* mosquitoes. Not all *Aedes* species spread Zika virus. At this time, we don't know if there are other non-*Aedes* mosquito species that could spread Zika virus.
- Zika virus is primarily spread through the bite of an Aedes aegypti or Aedes albopictus mosquito.
 - Aedes aegypti mosquitoes live in tropical, subtropical, and in some temperate climates. They are the
 primary vector of Zika, dengue, chikungunya, and other arboviral diseases. Because Aedes aegypti
 mosquitoes live near and prefer to feed on people, they are considered highly efficient at spreading these
 diseases.
 - Aedes albopictus mosquitoes live in tropical, subtropical, and temperate climates. They have adapted to survive in a broader temperature range and at cooler temperatures than Aedes aegypti. Because these mosquitoes feed on people and animals, they are less likely to spread viruses like Zika, dengue, or chikungunya. The strain of Ae. albopictus in the United States came from northern Japan in 1985 and is capable of living in more temperate climates.
- To produce eggs, the female mosquito bites people to feed on blood. When feeding, a mosquito will pierce the skin (like a needle) and inject saliva into a person's skin. This allows the disease-causing germ (for example, the Zika virus) into the site.
- In addition to Zika, the most common viruses and parasites spread through mosquito bites are:
 - Chikungunya
 - Dengue
 - Eastern equine encephalitis
 - Filariasis, including dirofilaria, which causes dog heartworm

- Jamestown Canyon virus disease
- Japanese encephalitis
- LaCrosse encephalitis
- o Malaria
- Rift Valley fever
- Ross River virus disease
- St. Louis encephalitis
- Venezuelan equine encephalitis
- Western equine encephalitis
- Yellow fever
- Once a mosquito is infected with Zika virus, it will remain infected for life. A mosquito lifespan is up to 30 days. There is no evidence that a mosquito infected with Zika will have a shorter than expected lifespan.
- Spread of Zika virus from an infected female mosquito to her eggs has not been well studied. One study shows that an adult female mosquito may pass Zika virus to 1 out of about 290 eggs.
 - Researchers at UTMB infected adult female Aedes aegypti and Aedes albopictus mosquitoes in the laboratory to see if Zika virus could be passed from a female to her eggs.
 - Zika virus was found to be passed to Aedes aegypti eggs.
 - From the data, transmission of Zika virus from an adult female mosquito to her eggs occurred in approximately 1 of 290 eggs. At this rate, an infected adult female might lay 2-3 infected eggs in her lifetime.
 - o In nature, we expect that the rate of transmission from a mosquito to egg would be less.
- Changes in the environment caused by climate change can influence the spread of mosquitoes.
 - These changes can affect
 - How quickly viruses replicate in mosquitoes
 - The life cycle of mosquitoes

the United States.

- The distribution of viruses, mosquitoes, or animal hosts
- Natural disasters in the continental United States have rarely been accompanied by outbreaks of viruses spread by mosquitoes. Flooding immediately washes away young mosquitoes, making mosquito populations temporarily decrease.
 - Following the disaster, mosquito eggs hatch and develop and mosquito populations surge (this takes about a week). New adult mosquitoes are not infected with virus until they bite an infected person or animal.
 - However, following flooding, mosquito eggs hatch and the mosquito population will increase.
 - Studies show that nuisance mosquitoes, not mosquitoes that typically spread viruses, will likely be a big problem after flooding.
 Post-flooding mosquito-borne diseases like Zika or West Nile are <u>unlikely</u> to be a large problem in
- Mosquitoes cannot survive high winds; they dry out and die. There is no evidence that high winds can successfully carry mosquitoes into new areas where they will survive.
- Mosquito eggs can survive drying out for up to 8 months. In the far south, like southern Florida and Texas, mosquitoes can be active year round. In those areas that stay warm year round, their abundance is driven by wet/dry seasons and not as much by temperature. In South Florida, the rainy season ends around the end of October/early November.
- Flies do not spread Zika. Only a small number of fly species will bite people. When a fly bites, it creates a wound and laps blood up from the site. When a fly bites, it does not directly inject saliva into the bite site like a mosquito does.
 - o Flies spread some diseases but fewer germs than mosquitoes because their feeding habits are different.

MODIFIED MOSQUITOES

Though their role in mosquito control has not yet been determined, CDC sees the use of genetically modified (GM)
mosquitoes and mosquitoes infected with Wolbachia (bacteria) as two promising new options for controlling
mosquitoes that can spread viruses like dengue, chikungunya, and Zika.

- Use of GM mosquitoes or *Wolbachia*-infected mosquitoes requires a special facility for rearing mosquitoes. Until the facility is built and operating, these mosquitoes may not be quickly available for mosquito control during an outbreak.
- GM or Wolbachia-infected mosquitoes must be released in large numbers multiple times in a community during mosquito season to decrease the overall mosquito population.
 - Releases are more successful if timed at the beginning of the mosquito season.
 - It takes at least 4-6 weeks to notice a reduction in local mosquito populations.
- GM mosquitoes and *Wolbachia*-infected mosquitoes are designed to reduce the overall mosquito population. These mosquitoes are designed to interrupt the mosquito life cycle by preventing the next generation of mosquitoes from surviving to become adults.
- Use of GM mosquitoes or *Wolbachia*-infected mosquitoes cannot and should not be used to replace traditional integrated mosquito management methods such as
 - Mosquito surveillance
 - Control of adult and young (larvae and pupae) mosquitoes
 - o Insecticide resistance monitoring
 - Personal protection (people protecting themselves from mosquito bites)
- In an outbreak, use of insecticides may still be a priority to keep people from getting infected. It is more important to kill infected adult mosquitoes that are spreading viruses immediately. Use of GM mosquitoes or *Wolbachia*-infected mosquitoes may not work quickly enough to stop an outbreak.
- There are no data to link GM mosquitoes released by Oxitec and the Zika outbreak or cases of microcephaly in Brazil. Oxitec released mosquitoes in only a few towns in Brazil. Occurrence of the Zika outbreak and cases of microcephaly have been reported from most states in Brazil.
 - Before Oxitec could release genetically modified mosquitoes in communities, the Brazilian government had to approve. These genetically modified mosquitoes have not been associated with or expected to cause any harmful effects in people.
- The US Food and Drug Administration (FDA) released its final environmental assessment on the Oxitec mosquito on August 5, 2016. FDA's update states that the FDA has completed the environmental review for a proposed field trial to determine whether the release of Oxitec Ltd.'s genetically engineered (GE) mosquitoes (OX513A) will suppress the local *Aedes aegypti* mosquito population in the release area at Key Haven, Florida. After considering thousands of public comments, the FDA has published a final environmental assessment (EA) and finding of no significant impact (FONSI) that agrees with the EA's conclusions that the proposed field trial will not have significant impacts on the environment. FDA's finalization of the EA and FONSI does not mean that Oxitec's GE mosquitoes are approved for commercial use. Oxitec is responsible for ensuring that all other local, state, and federal requirements are met before conducting the proposed field trial, and, together with its local partner, the Florida Keys Mosquito Control District, for determining whether and when to begin the proposed field trial in Key Haven, Florida.
 - The Final Environmental Assessment on Oxitec mosquitoes
 - o Additional information posted to the FDA website
 - Open field trials of Oxitec's genetically engineered mosquito have been conducted in Brazil, the Cayman Islands, Panama, and Malaysia.
 - Researchers have observed suppression of the targeted mosquito populations. They have not detected any adverse environmental or health outcomes.

SEXUAL TRANSMISSION

- Zika can be passed through sex from a person who has Zika to his or her sex partners.
- Sexual exposure includes sex without a condom, with a person who traveled to, or lives in an area with Zika, even if that person does not have symptoms of Zika.
 - o This includes vaginal, anal, and oral sex, and the sharing of sex toys.
- Zika can be passed through sex, even if the infected person does not have symptoms at the time.
 - o It can be passed from a person with Zika before their symptoms start, while they have symptoms, and after their symptoms end.
 - o The virus may also be passed by a person who carries the virus but never develops symptoms.
- Studies are underway to find out how long Zika stays in the semen and vaginal fluids of people who have Zika and how long it can be passed to sex partners. We know that Zika can remain in semen longer than in other body fluids, including vaginal fluids, urine, and blood.
- Zika has been found in genital fluids, including semen and vaginal fluids.
 - o Infectious, live Zika virus (virus that can be spread to others) has been found in semen at least 24 days after symptoms began.
 - Pieces of Zika virus (Zika RNA) have been found in semen as many as 188 days after symptoms began, and in vaginal and cervical fluids up to 3 and 11 days after symptoms began, respectively.
- Zika RNA may indicate the presence of live virus (which can be infectious to others), or it may simply indicate
 leftover genetic material (dead virus) that is no longer infectious to others. Finding virus RNA does not necessarily
 mean live virus is present or that a person can spread it to others.
 - o In most cases reported to date, no follow up testing was done to determine when infected men no longer had Zika virus in their semen.
 - In one case, sexual transmission is estimated to have occurred 32-41 days after onset of the man's symptoms.
 - CDC and other public health partners continue to study Zika virus and how it is spread and will share new
 information as it becomes available. This continuing research may help us find out
 - How long Zika can stay in genital fluids.
 - How common it is for Zika to be passed during sex by a man or woman.
 - If Zika can be passed through saliva during deep kissing.
 - If Zika passed to a pregnant woman during sex has a different risk for birth defects than Zika transmitted by a mosquito bite.
- Pregnant women with possible sexual exposure to Zika virus should be tested for Zika infection. Sexual exposure is defined as sex without a condom with someone who lives in or traveled to an area with Zika.
- CDC also recommends Zika virus <u>testing</u> for people who may have been exposed to Zika through sex **and** who have Zika <u>symptoms</u>.
- Testing blood, semen, vaginal fluids, or urine is not recommended to determine how likely a person is to pass Zika virus through sex. Because Zika virus can remain in semen longer than blood, someone might have a negative blood test but a positive semen test. The results of the tests are difficult to interpret.
 - Studies are underway to better understand the performance of these tests and how best to interpret the
 results. As additional information becomes available, CDC will share it.
- Available tests may not accurately identify the presence of Zika or a person's risk of passing it on through sex. As
 we learn more and as tests improve, these tests may become more helpful for determining a person's risk of
 passing Zika through sex.

[Refer to <u>Prevention</u> section for recommendations on preventing sexual transmission of Zika]

BREASTFEEDING

• There are currently no reports of transmission of Zika virus infection through breastfeeding.

- o Zika virus has been detected in breast milk.
- Based on available evidence, the benefits of breastfeeding outweigh any possible risk.
- Because of the benefits of breastfeeding, mothers are encouraged to breastfeed even in areas where Zika virus is found.
- CDC and the World Health Organization recommend that infants born to women with suspected, probable, or confirmed Zika virus infection, or who live in or have traveled to areas of Zika, should be fed according to normal infant feeding guidelines.
 - These infants should start breastfeeding within one hour of birth, be exclusively breastfed for 6 months, and have introduction of adequate, safe, and properly fed complementary foods, while continuing breastfeeding up to 2 years old or beyond.
 - o All mothers who decide to breastfeed should receive skilled support to initiate and sustain breastfeeding.
 - Mothers and families of infants born with congenital anomalies, such as microcephaly, or those
 presenting with feeding difficulties, should receive skilled feeding support from health professionals.
 - Multidisciplinary teams may be necessary for infants who need specialist support in infant feeding, which
 may be the case in particular for infants born with congenital anomalies, including microcephaly, and
 long-term management may be necessary.

BLOOD TRANSFUSION

- There is a strong possibility that Zika virus can be spread through <u>blood transfusions</u>.
 - Because many people infected with Zika virus don't have any symptoms, blood donors may not know they
 have been infected.
 - There have been suspected cases of Zika transmission through blood transfusion in Brazil. These reports
 are currently being investigated. During the Zika virus outbreak in French Polynesia in 2013-2014, 2.8% of
 blood donors tested positive for Zika. In previous outbreaks elsewhere, the virus has also been found in
 blood donors.
- Zika virus currently poses a low risk to the blood supply in the continental United States, but this could change depending on how many people become infected with the virus.
- To date, there have been no confirmed blood transfusion-transmission cases in the United States.

ZIKA VIRUS BLOOD SCREENING

- In areas of active transmission, the Food and Drug Administration (FDA) recommends that blood either be screened by laboratory testing, subjected to pathogen reduction technology (PRT), or outsourced from other areas. Blood donations that test positive for Zika virus are removed from the blood supply.
- To protect the US blood supply, CDC, in collaboration with the FDA, defines areas of active Zika virus transmission
 as having two or more locally acquired cases of Zika virus infection within 45 days. These defined areas of risk can
 be different from areas for which CDC has issued travel guidance, because of concerns about potential risk for
 blood safety.
 - The following are areas of active transmission of Zika virus in the continental United States for the purpose of blood and tissue safety intervention:
 - Miami-Dade County, FL
 - Palm Beach County, FL
- On August 26, 2016, FDA issued <u>revised guidance</u> to prevent the spread of Zika virus through the blood supply.
 This new FDA guidance calls for blood collection centers in the United States to screen all donated blood for Zika virus.

This guidance will be implemented immediately in Miami-Dade County and Palm Beach County, FL, in high-risk states within 4 weeks, and in all states within 12 weeks.

- To date, one donation tested positive during blood donor screening in Florida; a few other suspect positive donations are under investigation. These Zika-positive donations likely reflect infections associated with travel to an area of active Zika virus transmission. Identifying a positive blood donation shows that the screening is working and is preventing Zika from entering the blood supply.
- Blood donation centers routinely ask donors a set of standard questions before they donate blood. The questions
 help determine if donors are in good health and free of any diseases that could be transmitted by blood
 transfusion. If the donor's answers indicate they are not well or are at risk for having a disease transmissible by
 blood transfusion, they are not allowed to donate blood.
 - Blood donor screening on the basis of a questionnaire, without a laboratory test, is insufficient for identifying Zika-infected donors in areas with active mosquito-borne transmission of Zika virus due to the high rate of asymptomatic infection.
- Although there is no FDA-licensed test for Zika virus, on April 3, 2016 (Roche Molecular Systems, Inc.) and June 20, 2016 (Hologic, Inc. and Grifols), testing for Zika became available through two separate Investigational New Drug (IND) applications for blood collected in Puerto Rico and the continental United States.
 - Puerto Rico began using the Roche IND on April 3, 2016.
 - o Texas, Florida, Alabama, Mississippi, Georgia, and South Carolina are currently using one of the two INDs.
 - Additional locations in the US are expected to implement testing in the coming months.

PUERTO RICO SURVEY OF BLOOD COLLECTION CENTERS

- The Puerto Rico survey of blood collection centers was conducted February 10-24, 2016.
- The results of this survey were used to guide a federally supported coordinated effort to address the blood supply and safety challenges in Puerto Rico. This effort included importing all blood components from the continental United States at a volume sufficient to meet the demand projected from the 2015 estimates, beginning March 5, 2016, until a nucleic acid screening test was implemented under IND protocol beginning April 4, 2016.
- Efforts to implement PRT for apheresis platelets and plasma collections in Puerto Rico are currently under way, and evaluation trials to determine safety and efficacy of investigational PRT for red blood cells (RBCs) are in planning stages.

RISK

- Anyone who lives in or travels to an area where Zika virus is found and has not already been infected with Zika virus can get it from mosquito bites.
- A person with Zika virus can pass it to his or her sex partners during sex. See <u>Sexual Transmission</u> and <u>Preventing</u> Sexual Transmission.
- Based on information about similar infections, once you have been infected with Zika virus, you are likely to be protected from future infections.

PREVENTION

- There is no vaccine to prevent Zika virus disease.
- The best way to prevent diseases spread by mosquitoes is to protect yourself and your family from mosquito bites.
 - Wear long-sleeved shirts and long pants.
 - Stay in places with air conditioning and window and door screens to keep mosquitoes outside.
 - Treat your clothing and gear with <u>permethrin</u> or buy pre-treated items.
 - Use Environmental Protection Agency (EPA)-registered insect repellents. See Insect Repellent section.

 Sleep under a mosquito bed net if air conditioned or screened rooms are not available or if sleeping outdoors.

- During the first week of infection, Zika virus can be found in a person's blood and can pass from an infected person to a mosquito through mosquito bites. An infected mosquito can then spread the virus to other people.
 - To help prevent others from getting sick, strictly follow steps to prevent mosquito bites during the first week of illness.
- Even if they do not feel sick, travelers returning to the United States from an area with Zika should take steps to prevent mosquito bites for 3 weeks. These steps will prevent them from passing Zika to mosquitoes that could spread the virus to other people.
- If you have a baby or child
 - o Do not use insect repellent on babies younger than 2 months old.
 - Do not use products containing oil of lemon eucalyptus or para-menthane-diol on children younger than 3
 years old.
 - Dress your child in clothing that covers arms and legs.
 - o Cover crib, stroller, and baby carrier with mosquito netting.
 - o Do not apply insect repellent onto a child's hands, eyes, mouth, and cut or irritated skin.
 - o Adults: Spray insect repellent onto your hands and then apply to a child's face.

INSECT REPELLENT

- CDC recommends using <u>EPA-registered insect repellents</u> with one of the following active ingredients: DEET, picaridin, IR3535, or oil of lemon eucalyptus or para-menthane-diol.
 - o Choosing an EPA-registered repellent ensures the EPA has evaluated the product for effectiveness.
- Insect repellents registered by the EPA repel the mosquitoes that spread Zika and other viruses like dengue, chikungunya, and West Nile.
- When used as directed, EPA-registered insect repellents are proven safe and effective even for pregnant and breastfeeding women.
- Always follow the product label instructions.
- Reapply insect repellent as directed.
- Do not spray repellent on the skin under clothing.
- If you are also using sunscreen, apply sunscreen before applying insect repellent.
- Treat clothing and gear with permethrin or buy permethrin-treated items.
 - The EPA has reviewed scientific studies on the use of permethrin-treated clothing. Based on EPA's review, there is no evidence of reproductive or developmental effects to mother or child following exposure to permethrin.
 - Treated clothing remains protective after multiple washings. See product information to learn how long the protection will last.
 - o If treating items yourself, follow the product instructions carefully.
 - o Do NOT use permethrin products directly on skin. They are intended to treat clothing.
 - In some places, such as Puerto Rico, where permethrin products have been used for years in mosquito control efforts, mosquitoes have become resistant to it. In areas with high levels of resistance, use of permethrin and related products is not likely to be effective. Contact local authorities or a mosquito control district for more information on pesticides.
- We do not know the effectiveness of non-EPA registered insect repellents, including some natural repellents.
 - Some natural insect repellents, often made with natural oils, have not been tested for effectiveness.
 Homemade insect repellents may not protect you from mosquito bites.
- Some natural products are EPA-registered.
 - o These natural products with EPA registration include para-menthane-diol and oil of lemon eucalyptus.

- For children
 - o Do not use insect repellents on babies younger than 2 months old.
 - Mosquito netting can be used to cover babies younger than 2 months old in carriers, strollers, or cribs to protect them from mosquito bites.
 - Do not use products containing oil of lemon eucalyptus or para-menthane-diol on children younger than 3
 years old.
 - o Do not apply insect repellent onto a child's hands, eyes, mouth, and cut or irritated skin.
 - o Adults: Spray insect repellent onto your hands and then apply to a child's face.

CONTROLLING MOSQUITOES AT HOME

- To control mosquitoes outside your home:
 - Once a week, empty and scrub, turn over, cover, or throw out any items that hold water like tires, buckets, planters, toys, pools, birdbaths, flowerpot saucers, or trash containers. Mosquitoes lay eggs near water.
 - Tightly cover water storage containers (buckets, cisterns, rain barrels) so that mosquitoes cannot get inside to lay eggs.
 - For containers without lids, use wire mesh with holes smaller than an adult mosquito.
 - Use larvicides to kill young mosquitoes in large containers of water that will not be used for drinking and cannot be covered or dumped out.
 - o **Use an outdoor insect spray** made to kill mosquitoes in areas where they rest.
 - Mosquitoes rest in dark, humid areas like under patio furniture, or under the carport or garage.
 - o **If you have a septic tank,** repair cracks or gaps. Cover open vent or plumbing pipes. Use wire mesh with holes smaller than an adult mosquito.
- To control mosquitoes inside your home:
 - o **Install or repair and use window and door screens**. Do not leave doors propped open.
 - Use air conditioning when possible.
 - Once a week, empty and scrub, turn over, cover, or throw out any items that hold water like vases and flowerpot saucers. Mosquitoes lay eggs near water.
 - Kill mosquitoes inside your home. Use an indoor insect fogger* or indoor insect spray* to kill mosquitoes and treat areas where they rest. These products work immediately, and may need to be reapplied. When using insecticides, always follow label directions. Only using insecticide will not keep your home free of mosquitoes.
 - Mosquitoes rest in dark, humid places like under the sink, in closets, under furniture, or in the laundry room.
- Do-it-yourself products*:

Product	Active ingredient	Brand name examples**	How long it works
Indoor insect spray	Imidacloprid, β-	Home Pest Insect Killer, Raid,	7-10 days
	Cyfluthrin	Ortho, HotShot, EcoLogic	
Indoor insect fogger	Tetramethrin,	Hot Shot, Raid, Real Kill,	Up to 6 weeks
	Cypermethrin	Spectracide	

^{**}Insecticide brand names are provided for your information only. The Centers for Disease Control and Prevention and the U.S. Department of Health and Human Services cannot recommend or endorse any name brand products.

PREVENTING SEXUAL TRANSMISSION

- Condoms can reduce the chance of getting Zika from sex.
 - Condoms include male and female condoms.

o To be effective, <u>condoms</u> should be used from start to finish, every time during vaginal, anal, and oral sex and the sharing of sex toys.

- Dental dams (latex or polyurethane sheets) may also be used for certain types of oral sex (mouth to vagina or mouth to anus).
- Not sharing sex toys may also reduce the risk of spreading Zika to sex partners.
- Not having sex eliminates the risk of getting Zika from sex.
- The risk of Zika from sex is of greatest concern for pregnant women, who can pass Zika to their developing fetus if infected during pregnancy. Because Zika can cause birth defects, pregnant women with partners who live in or traveled to an area with Zika should protect themselves throughout their entire pregnancy.
- Pregnant couples with a partner who lives in or traveled to an area with Zika should:
 - Use condoms every time they have sex or not have sex during the pregnancy. This is important, even if the pregnant woman's partner does not have symptoms of Zika or feel sick.
 - Not share sex toys throughout the entire pregnancy.
 - o This applies to couples in which one or both partners have traveled to or live in an area with Zika.
- Pregnant couples who are concerned that one of them may have Zika should:
 - Tell their healthcare provider immediately about
 - Symptoms of Zika
 - Each partner's travel history
 - How long either partner stayed in an area with Zika
 - If either partner took steps to prevent mosquito bites while in an area with Zika
 - If they had sex without a condom
- **Couples who are considering pregnancy** should talk to their healthcare provider if one or both partners live in or traveled to an area with Zika.
- Anyone who is not pregnant or trying to get pregnant who wants to avoid getting or passing Zika during sex can use condoms every time they have sex or not have sex. The recommended period of time for taking these precautions will depend on the person's or couple's situation.
 - Couples with a partner who lives in or has <u>traveled</u> to an <u>area with Zika</u> should use condoms or not have sex for:
 - At least 8 weeks after a Zika diagnosis or start of symptoms if the traveling partner is female.
 - At least 6 months after a Zika diagnosis or start of symptoms if the traveling partner is male. This long period is because Zika stays in semen longer than in other body fluids.
 - At least 8 weeks after returning if the traveling partner (male or female) has no symptoms.
 - Couples living in an <u>area with Zika</u> can use condoms or not have sex. If either partner develops symptoms
 of Zika or has concerns, they should talk to a healthcare provider.
 - Couples who are considering condoms or abstinence should weigh the personal risks and benefits, including
 - The mild nature of the illness for many people
 - Either partner's exposure to mosquitoes while in an area with Zika
 - Plans for pregnancy (if appropriate) and access to birth control
 - Access to condoms
 - Desire for intimacy, including willingness to use condoms or not have sex
 - Ability to use condoms or not have sex
 - In addition to condoms, couples who do not want to get pregnant should use the most effective birth control methods that they can use correctly and consistently. They should talk to their doctor about available options.
- Recommendations for condom use among pregnant couples do not change if a person with possible Zika virus
 exposure tests negative for Zika virus infection. Pregnant couples in which one or both partners have traveled to or

live in an area with Zika should use a condom from start to finish, every time they have sex or should not have sex for the entire pregnancy. To be effective, <u>condoms</u> must be used from start to finish, every time you have sex. This includes vaginal, anal, and oral sex and the sharing of sex toys. Condoms include male and female condoms.

PYRIPROXYFEN

- Several media reports in February 2016 suggested that a larvicide called pyriproxyfen might be linked with
 microcephaly. These media reports appear to be based on a February 3 publication authored by an Argentine
 physicians' organization, which claims that the use of pyriproxyfen in drinking water in Brazil is responsible for the
 country's increase in microcephaly cases.
- The World Health Organization (WHO) has approved the use of pyriproxyfen for the control of disease-carrying mosquitoes.
- Pyriproxyfen is a registered larvicide in Brazil and other countries, it has been used for decades, and it has not been linked with microcephaly.
- Exposure to pyriproxyfen would not explain recent study results showing the presence of Zika virus in the brains of infants born with microcephaly.
- In a published study, CDC scientists concluded that Zika virus infection during pregnancy is a cause of microcephaly and other severe fetal brain defects. Scientists continue to study other potential health problems that Zika virus infection during pregnancy may cause.

AERIAL SPRAYING

- Aerial spraying uses airplanes to spray large areas with small amounts of insecticide. This type of spraying is safe, quick, and efficient.
- Aerial spraying is the preferred method for applying insecticide when people in a large area are getting sick with
 viruses mosquitoes can carry like Zika or West Nile (or dengue and chikungunya throughout the US territories) or
 when large numbers of infected mosquitoes are found.
 - o It helps control and immediately reduce the number of mosquitoes that can spread viruses, like Zika.
- Aerial spraying has been used successfully for decades in the United States and its territories to help control and immediately reduce the number of mosquitoes that can spread viruses, like Zika, dengue, chikungunya and West Nile viruses.
- Airplanes spray insecticide that kills either young or adult mosquitoes. These products are called larvicides (to kill young mosquitoes) and adulticides (to kill adult mosquitoes).
- When a mosquito control district decides to spray wide areas of a community, it must use an EPA-registered product in accordance with label instructions, and a licensed professional must apply it.
 - o Local government or mosquito control program will decide which type of insecticide to use.
- Aerial spraying occurs when mosquitoes are active, sometime between the early evening, close to sunset, and the early morning, close to sunrise.
 - This is when most insects, including honey bees, are not active, making them less likely to be affected by spraying.
 - However, adulticide sprays may kill other insects that come in contact with the spray.
 - Larvicides sprays will not kill other insects.
- During aerial spraying, a very small amount of insecticide is sprayed over the area, about 1 ounce (2 tablespoons) per acre, or about the size of some soccer fields.
 - o This small amount does not pose a health risk to people or pets in the area that is sprayed.
 - You do not need to leave an area when it is sprayed. You are unlikely to breathe in or touch anything that
 has enough insecticide on it to cause health problems.

 Aerial spraying of adulticides and larvicides will not cause long-term harm to the environment or local ecosystems, even if spraying is repeated.

TREATMENT

- There is no specific medicine or vaccine for Zika virus.
- Treat the symptoms.
 - o Get plenty of rest.
 - Drink fluids to prevent dehydration.
 - o Take medicine such as acetaminophen (Tylenol®) to reduce fever and pain.
 - o Do not take aspirin or other non-steroidal anti-inflammatory drugs (NSAIDS) until dengue can be ruled out to reduce the risk of bleeding.
 - o If you are taking medicine for another medical condition, talk to your doctor or other healthcare provider before taking additional medication.
- During the first week of infection, Zika virus can be found in a person's blood. The virus can be passed from an infected person to a mosquito through mosquito bites. An infected mosquito can then spread the virus to other people.
 - To help prevent others from getting sick, strictly <u>follow steps to prevent mosquito bites</u> during the first week of illness.

ZIKA & PREGNANCY

- Because of the potential risks of Zika virus infection during pregnancy, CDC's top priority for the Zika response is to
 protect pregnant women and their fetuses.
- Zika virus can pass from a pregnant woman to her fetus during pregnancy or around the time of birth. We do not know how often this happens.
- Zika virus infection during pregnancy can cause <u>microcephaly</u> and other severe fetal brain defects. Scientists are investigating other potential health problems that Zika virus infection during pregnancy may cause.
 - See Microcephaly section.
- In addition to microcephaly, doctors have found other problems in pregnancies and among fetuses and infants infected with Zika virus before birth, such as miscarriage, stillbirth, absent or poorly developed brain structures, defects of the eye, hearing deficits, and impaired growth. Although Zika virus has been linked with these other problems in infants, there is more to learn. Researchers are collecting data to better understand the extent of Zika virus' impact on mothers and their children.
- We expect that pregnant women who develop Zika will have similar illness to people who are not pregnant. No
 evidence exists to suggest that pregnant women are more susceptible or experience more severe disease during
 pregnancy.
- We do not know if pregnant women are more likely to develop <u>symptoms</u> compared to the general population.
- We do not know if pregnant women are more likely to get <u>Guillain-Barré syndrome</u> if infected with Zika.
- Pregnant women should talk to a doctor or other healthcare provider if they or their sex partners recently traveled to an area with Zika, even if they don't feel sick.
- Pregnant women should see a doctor or other healthcare provider if they develop a fever, rash, joint pain, or red eyes during their trip or within 2 weeks after traveling to a country where Zika has been reported. They should tell the doctor or other healthcare provider where they traveled.
- Because Zika infection is a cause of microcephaly, pregnant women should strictly follow steps to <u>prevent mosquito bites</u> and to <u>protect against sexual transmission</u>.
 - See Prevention and Preventing Sexual Transmission.

For guidance on suggested timeframes for delaying pregnancy, see <u>Couples Interested in Conceiving</u>.

MICROCEPHALY

- Zika can cause microcephaly, a severe birth defect that is a sign of a problem with brain development.
- Microcephaly is a condition in which a baby's head is much smaller than expected. During pregnancy, a baby's
 head grows because the baby's brain grows. Microcephaly can occur because a baby's brain has not developed
 properly during pregnancy or has stopped growing after birth.
- CDC concluded that Zika virus infection is a cause of microcephaly.
 - More lab testing and other studies are planned to learn more about the risks of Zika virus infection during pregnancy.
 - CDC scientists announced that enough evidence has accumulated to conclude that Zika virus infection during pregnancy is a cause of microcephaly and other severe fetal brain defects.
 - Recognizing that Zika is a cause of certain birth defects does not mean that every pregnant woman
 infected with Zika will have a baby with a birth defect. It means that infection with Zika during pregnancy
 increases the chances for these problems.
 - Although studies to date have linked Zika with certain birth defects or other pregnancy problems, it's important to remember that even in places with active Zika transmission, women are delivering infants that appear to be healthy.
- We do not know if a newborn who gets Zika at birth will develop microcephaly after birth, which is called acquired
 microcephaly.
 - o Babies can acquire microcephaly if their head growth slows or fails to develop after birth.
 - There have been no reports of Zika infection around the time of birth and acquired microcephaly.
 - All reports of microcephaly so far have been congenital microcephaly, meaning the microcephaly occurred before birth.
- Currently, there is no evidence to suggest that past Zika virus infection poses an increased risk of birth defects for future pregnancies once the virus has cleared the body.
- Zika virus usually remains in the blood of an infected person for about a week. There is no evidence that the virus will cause infection in a pregnancy that occurs after the virus is cleared from the mother's blood. The virus can be present in semen and urine longer than in blood.
- Babies with microcephaly can have a range of other health problems, depending on how severe their microcephaly
 is. These problems can range from mild to severe and are often lifelong. In some cases, these problems can be lifethreatening.
 - Seizures
 - Developmental delay, such as problems with speech or other developmental milestones (like sitting, standing, and walking)
 - o Intellectual disability (decreased ability to learn and function in daily life)
 - o Problems with movement and balance
 - Feeding problems, such as difficulty swallowing
 - Hearing loss
 - Vision problems
- Because it is difficult to predict at birth what problems babies will have from microcephaly, they often need close follow-up through regular check-ups with a doctor or other healthcare provider to track their growth and development.

• A new analysis from CDC estimates that 5,900-10,300 pregnant women might be infected during the Zika virus outbreak in Puerto Rico in 2016.

- Because the full spectrum of health effects that might occur from Zika virus infection during pregnancy is unknown, this range reflects the number of infants at risk for health problems due to Zika.
- CDC scientists estimated that 100-270 of these infections in pregnant women might lead to microcephaly in infants between mid-2016 and mid-2017.
- These estimates underscore the urgent actions being taken in Puerto Rico to prevent Zika and plan for services for families affected by Zika.
- Strategies to prevent Zika include mosquito control programs, dissemination of Zika Prevention Kits (containing health information, insect repellent, a bed net, standing water treatment tablets, and condoms), and approaches to improve access to effective birth control for women and their partners who want to avoid or delay pregnancy.
- In addition to these prevention efforts, Puerto Rico and CDC are monitoring Zika infections in pregnant women, and the outcomes of these pregnancies, to link the mothers and their infants to medical specialists and support services if needed.

DIAGNOSIS

- During pregnancy, microcephaly can sometimes be diagnosed during an ultrasound (which creates pictures of the baby). Multiple ultrasounds may be needed to detect an abnormality.
- Microcephaly might not be detectable until late in the second or early in the third trimester of pregnancy.
- CDC has interim guidelines for testing and evaluating an infant with possible congenital Zika virus infection.
 - The accuracy of the test varies depending on the type of test, the timing of the test during pregnancy, the specific equipment used for the test, and the person conducting the test.
- CDC provides information on diagnosing birth defects both during pregnancy and after birth. Some tests need to
 be done during a particular time in pregnancy, but others such as an ultrasound can be done at any time during
 pregnancy. High resolution ultrasounds (also known as Level 2 ultrasounds) are used to look in more detail for
 possible birth defects and are usually completed between weeks 18 and 22 of pregnancy. They can also be
 performed later if risks are detected later in pregnancy.

CONTRACEPTION TO PREVENT UNINTENDED PREGNANCY DURING THE ZIKA VIRUS OUTBREAK

- Preventing unintended pregnancy during the Zika virus outbreak is a primary strategy to reduce the number of pregnancies affected by Zika virus.
- Despite the availability of a wide range of FDA-approved contraceptives, unintended pregnancy, or a pregnancy that is mistimed or unwanted, remains common in the United States. Nearly half of all pregnancies in the United States are unintended.
- The best way for sexually active women and their partners to reduce the risk of unintended pregnancy is to use effective birth control consistently and correctly.
- Contraception should be considered as a medical countermeasure to unintended pregnancy, reducing the effects of Zika virus by preventing potential poor birth outcomes associated with infection.
- Contraceptive use varies across states.
- For states and jurisdictions with possible Zika virus transmission, CDC recommends that healthcare providers
 counsel women and their partners who do not want to become pregnant during this Zika virus outbreak.
- Decisions about pregnancy planning are personal and circumstances for women and their partners are different.
 Information about Zika from a healthcare provider may be helpful in considering whether or not to become pregnant.
- Prior to the 2016 Zika virus outbreak,

 Less than 1 in 4 sexually active women of reproductive age and fewer than one in 10 sexually active female high school students reported using long-acting reversible contraception (LARC).

- Postpartum women reported highest use of LARC. One in three women who recently delivered a live birth reported using LARC.
- Moderately effective and less effective contraceptive methods, including pills, patches, rings, injections, condoms and other barrier methods, were used more frequently than highly effective methods across all age groups and race/ethnicity.
- A substantial proportion of sexually active female high school students and women at risk for unintended pregnancy were not using contraception, or were using less effective methods.

RECOMMENDATIONS FOR COUPLES INTERESTED IN CONCEIVING

- CDC issued new guidance and information to prevent Zika virus transmission and negative health outcomes, including <u>updated interim guidance for healthcare professionals</u> for counseling patients about pregnancy planning and the timing of pregnancy after possible exposure to Zika virus.
 - For women and men who have been diagnosed with Zika virus disease or who have symptoms of Zika
 (fever, rash, joint pain, or red eyes) after possible exposure to Zika virus, CDC recommends that
 healthcare providers
 - Advise women to wait at least 8 weeks after their symptoms first appeared before trying to get pregnant.
 - Advise men to wait at least 6 months after their symptoms first appeared before trying to get their partner pregnant.
 - People with Zika should use condoms for sex or not have sex during this time period if they are concerned about the possibility of transmitting Zika virus to their sex partners. To be effective, condoms must be used every time, from start to finish during vaginal, anal, and oral sex and the sharing of sex toys.
 - For men and women without symptoms of Zika virus but who had possible exposure to Zika from recent travel or sexual contact, healthcare providers should recommend their patients wait at least 8 weeks after their possible exposure before trying to get pregnant.
 - Men and women without symptoms of Zika virus who live in an area with active Zika transmission should talk with their healthcare providers about their pregnancy plans during a Zika virus outbreak, the potential risks of Zika, and how they can prevent Zika virus infection during the pregnancy.
 - For healthcare professionals: Decisions about pregnancy planning are deeply personal and very complex.
 Circumstances for women and their partners will vary, and these circumstances are best discussed as part of clinical care for a woman and her partner during a visit with a trusted healthcare professional.

US ZIKA PREGNANCY REGISTRY

- CDC established the <u>US Zika Pregnancy Registry</u> and is collaborating with state, tribal, local, and territorial health
 departments to collect information about pregnancy and infant outcomes among pregnant women with laboratory
 evidence of Zika virus infection and their infants.
 - o CDC also developed a similar system, the Zika Active Pregnancy Surveillance System, in Puerto Rico.
- Health departments are working with healthcare providers to collect information that is needed to counsel pregnant women, and to plan for services to meet the needs of families affected by Zika. They are collecting information about exposure to Zika, the presence or absence of symptoms and pregnancy complications, prenatal Zika testing, pregnancy and birth outcomes, and infant health and development.
- The data collected through these registries will provide additional, more comprehensive information to complement notifiable disease case reporting and will be used to update recommendations for clinical care, to

plan for services for pregnant women and families affected by Zika virus, and to improve prevention of Zika virus infection during pregnancy.

• CDC maintains a 24/7 consultation service for health officials and healthcare providers caring for pregnant women. To contact the service, call 770-488-7100 or email ZIKAMCH@cdc.gov.

NATIONAL REPORTING: PREGNANT WOMEN & OUTCOMES

- Starting May 20, 2016, national reporting of the number of US pregnant women affected by Zika virus changed.
 - Before May 20, 2016, CDC reported the number of pregnant women who met both of the following criteria: 1) they had symptoms or pregnancy complications consistent with Zika virus disease and 2) they had laboratory test results that show evidence of a recent Zika virus infection.
 - Starting May 20, 2016, CDC began <u>reporting pregnancy data</u> from two enhanced surveillance systems:
 the US Zika Pregnancy Registry and the Puerto Rico Zika Active Pregnancy Surveillance System. Both of
 these systems include pregnant women with any laboratory evidence of possible Zika virus infection, with
 or without symptoms.
- These updated numbers reflect counts of pregnant women in the United States, including US territories, with any laboratory evidence of possible Zika virus infection, with or without symptoms or pregnancy complications.
 - Pregnant women with laboratory evidence of possible Zika virus infection include those in whom Zika virus particles have been detected and those with evidence of an immune reaction to a recent virus that may be possible Zika infection.
 - The registries cast a broad net to ensure that CDC is monitoring pregnancies at risk for poor outcomes associated with possible Zika infection.
- Given that the <u>US Zika Pregnancy Registry (USZPR)</u> and the Puerto Rico Zika Active Pregnancy Surveillance System (ZAPSS) aim to provide a complete and representative description of pregnancy and infant outcomes associated with Zika, participation by all jurisdictions is critical.
- This information will help healthcare providers as they counsel pregnant women affected by Zika and is essential for planning at the federal, state, and local levels for clinical, public health, and other services needed to support pregnant women and families affected by Zika.
- What these new numbers do not show
 - These new numbers are *not comparable* to the previous reports from ArboNET. These updated numbers reflect a different, broader population of pregnant women.
 - These updated numbers are not *real time* estimates. They will reflect the number of pregnant women reported with any laboratory evidence of possible Zika virus infection every Thursday the week prior; numbers will be delayed one week.
- This updated reporting aligns with recommendations for ongoing monitoring of pregnancies at risk for poor outcomes associated with Zika, based on scientists' current understanding of the effects of Zika infection during pregnancy.
- The registries will report two numbers that reflect the aggregated data from the US states and DC and separately the combined total from the US territories. CDC is not reporting individual state, tribal, territorial or jurisdictional level data. Comprehensive national information will facilitate and improve the public health response.
- Starting June 16, 2016, CDC began reporting poor outcomes of pregnancies with laboratory evidence of possible Zika virus infection. CDC will report two types of outcomes:
 - o Live-born infants with birth defects and
 - Pregnancy losses with birth defects
- The poor pregnancy outcomes reported include those that are known to be caused by Zika (e.g., microcephaly and other severe fetal brain defects) as well as others associated with Zika infection during pregnancy (e.g., eye defects, hearing abnormalities) that may be linked to Zika.

ZIKA & PREGNANCY IN COLOMBIA

• Findings from *New England Journal of Medicine* paper (June 15, 2016) entitled Zika Virus Disease in Colombia: Preliminary Report:

- Nearly 66,000 people, including nearly 12,000 pregnant women, were reported to have Zika virus disease
 in Colombia from August 9, 2015, through April 2, 2016.
- The reported rate of Zika virus disease was about two times higher in women than men overall and about three times higher in women aged 15-29 years compared to men of the same age.
 - This could be the result of true increased risk or the result of other factors like reporting/testing bias or increased healthcare-seeking behavior.
- Pregnant women infected with Zika virus during their first or second trimester were still pregnant at the time of this report (report cut-off date was May 2, 2016). Data on these pregnancy outcomes will be reported when available.
- Among a subset of the pregnant women with Zika virus disease, a majority (over 90%) of those infected in the third trimester delivered no infants with apparent birth defects, including microcephaly.
 - Although these preliminary data from Colombia suggest that Zika virus infection during the third trimester of pregnancy is not linked to birth defects like microcephaly, continued monitoring of the impact of Zika virus upon pregnancy and infant outcomes is ongoing.
- Cases of microcephaly are starting to emerge in Colombia.
 - From January 1 to April 28, 2016, four infants with microcephaly had laboratory evidence of congenital Zika infection, and all were born to women with asymptomatic Zika virus infections.
 - This suggests that poor pregnancy and infant outcomes like microcephaly can occur in women with Zika virus infection regardless of whether they have symptoms.
- o CDC and Colombia's Instituto Nacional de Salud will continue to collaborate to provide critical scientific information about Zika virus infection during pregnancy.

GUILLAIN-BARRÉ SYNDROME

- Several countries that have experienced Zika outbreaks recently have reported increases in people who have Guillain-Barré syndrome (GBS).
- Current CDC research suggests that GBS is strongly associated with Zika; however, only a small proportion of people with recent Zika virus infection get GBS. CDC is continuing to investigate the link between GBS and Zika to learn more.
- GBS is an uncommon illness of the nervous system in which a person's own immune system damages the nervous cells, causing muscle weakness, and sometimes, paralysis.
- GBS symptoms include weakness of the arms and legs and, in severe cases, can affect the muscles that control breathing.
- These symptoms can last a few weeks or several months. Most people fully recover from GBS, though some people have permanent damage. Very few people die from GBS.
- Researchers do not fully understand what causes GBS. Most people with GBS report an infection before they have GBS symptoms. Rarely, vaccination has also been associated with the onset of GBS (for example, the 1976 Swine influenza vaccine).
- An estimated 3,000 to 6,000 people, or 1-2 cases for every 100,000 people, develop GBS each year in the United States. Most cases of GBS occur for no known reason, and true "clusters" of cases of GBS are very unusual.
- If you want to know more about the number of GBS cases in a certain area, contact the state or local health department in the state where the cases happen. CDC collaborates with state and local health departments to investigate reports of possibly unusually large numbers or "clusters" of GBS cases.

TRAVEL RECOMMENDATIONS

- Travelers who go to places with outbreaks of Zika can be infected with Zika virus.
 - o Travelers can protect themselves by preventing mosquito bites.
- Some travelers become infected while traveling but do not get sick until they return home. Be aware of any illness or symptoms during your trip or after you return home. Tell your doctor or other healthcare provider where you have traveled and when you were there.
- Even if they do not feel sick, travelers returning from an area with Zika should take steps to prevent mosquito bites for 3 weeks so they do not spread Zika to uninfected mosquitoes.
- The mosquitoes that spread Zika usually do not live at elevations above 6,500 feet (2,000 meters). Travelers who plan to be only in areas above this elevation are at a very low risk of getting Zika from a mosquito.
- Sexual transmission of Zika virus is possible, so travelers are encouraged to use condoms or not have sex. See Preventing Sexual Transmission.
- Until more is known, CDC recommends the following:
 - Pregnant women
 - Should not travel to any area below 6,500 feet (2,000 meters) where Zika virus is spreading.
 - If you must travel to one of these areas, talk to your doctor or other healthcare provider first and strictly follow <u>steps to prevent mosquito bites</u> during the trip. If your itinerary is limited **entirely** to areas above 6,500 feet, there is minimal risk of getting Zika from a mosquito.
 - If your sex partner lives in or has traveled to an area where Zika is spreading, use a condom every time you have sex or do not have sex during the pregnancy. See <u>Preventing Sexual Transmission</u>.
 - Women trying to get pregnant
 - Before you or your partner travel, talk to your doctor or other healthcare provider about your plans to become pregnant and the risk of Zika virus infection.
 - You and your partner should strictly follow steps to prevent mosquito bites during the trip.
 - See <u>Recommendations for Couples Interested in Conceiving</u>.
- There are no restrictions for travelers entering the United States who have contracted Zika virus. CDC is not conducting enhanced entry screening of arriving travelers for Zika at this time.
 - Because many people who have Zika do not have symptoms, entry screening will not work to prevent imported cases. CDC and Customs and Border Protection are working together to assess the situation and determine necessary measures.
 - CDC has routine steps to detect sick travelers entering the United States, including requirements for ships and airplanes arriving in the United States to report certain illnesses to CDC. State and territorial health departments routinely notify CDC when cases of Zika are detected in the United States.

TRAVEL TO FLORIDA

- The Florida Department of Health has identified two areas of Miami-Dade County where Zika is being spread by
 mosquitoes. In addition to the previously identified area in the Wynwood neighborhood, there is now mosquitoborne spread of Zika virus in a section of Miami Beach.
- This guidance is for people who live in or traveled to the identified area of Miami Beach any time after July 14. This guidance also still applies for those who live in or traveled to the previously identified Wynwood area any time after June 15. These timeframes are based on the earliest time symptoms can start and the maximum 2-week incubation period for Zika virus.
 - o Pregnant women should not travel to these areas.
 - Pregnant women and their partners who are concerned about being exposed to Zika may want to consider postponing nonessential travel to all parts of Miami-Dade County.

 Pregnant women who live in or frequently travel to these areas should be tested in the first and second trimester of pregnancy.

- o Women and men who are thinking about getting pregnant and who live in or frequently travel to these areas should talk to their healthcare provider.
- See the full guidance <u>here</u>.

TRAVEL NOTICES

- CDC has issued <u>travel notices</u> (level 2 alert, "practice enhanced precautions") for people traveling to international destinations and overseas US territories where Zika virus is spreading.
 - o Specific areas where Zika is spreading are often difficult to determine and are likely to change over time.
 - As more information becomes available, CDC's <u>Zika travel notices</u> will be updated. Please check back frequently for the most up-to-date recommendations.
- CDC regularly issues level 2 alert travel notices when recommending special precautions for travelers because of a specific outbreak or situation.
 - Special precautions might mean getting a certain vaccine or taking a certain medicine that would not usually be recommended for that destination. Sometimes the special precaution is that a certain group should avoid travel.
- Countries and territories where Zika is spreading are included in the travel notice.
 - Local transmission means that mosquitoes in affected areas have been infected with Zika virus and are spreading it to people.
 - Countries and territories with imported cases are not included in the travel notice. Imported cases occur
 when people are infected with Zika during travel to an affected area and then return to their home
 countries.
 - Countries with past Zika transmission are not included. CDC has had Zika travel notices in the past for several other countries, but those were removed as outbreaks ended.
- CDC has posted notices that include maps that show elevation levels in countries with Zika.
- CDC has posted <u>maps</u> that show elevation levels in countries with Zika.
- The mosquitoes that spread Zika usually do not live at elevations above 6,500 feet (2,000 meters). Travelers whose itineraries are limited **entirely** to areas above this elevation are at minimal risk of getting Zika from a mosquito.

2016 OLYMPIC AND PARALYMPIC GAMES IN BRAZIL

- Brazil, along with many destinations in the Americas, is experiencing an outbreak of Zika virus.
- CDC has issued guidance for travel to the 2016 Summer Olympic and Paralympic Games in Brazil.
 - Guidance for travel to the Olympics includes information about travel vaccines, insect bite prevention, safe food and water practices, and safety and security in crowds.
 - Because Zika virus is spreading locally in Brazil and Zika virus infection in pregnant women can cause serious birth defects, CDC has special travel recommendations for pregnant women.
- Based on what we currently know, CDC does not recommend cancelling or postponing the 2016 Olympic and Paralympic Games.
 - o CDC is working closely with the US Olympic Committee and Brazilian health authorities, and will update our guidance if needed.
 - Travel to the Olympic Games is a small fraction of overall global travel to countries that have ongoing Zika transmission. Travel to the Olympic Games represents about 0.25% of total aviation travel annually to areas with Zika.

• The Games will occur during the winter months in Brazil. Although risk of mosquito-borne diseases is lower during winter, travelers should still take personal protective measures to prevent mosquito bites.

- CDC understands people travel for a variety of reasons: weddings, tourism, Olympics/Paralympics, etc. Our guidance for travel to Brazil and other areas with Zika remains:
 - o Pregnant women should not travel to Brazil for the games or to other areas with Zika.
 - After returning from the games (or any area with Zika), travelers with pregnant partners should use condoms or not have sex during the pregnancy.
 - All travelers can take steps to avoid Zika while at the games, and prevent spreading Zika when they get back to the U.S.
 - Prevent mosquito bites both during travel and for 3 weeks after returning to the U.S.
 - Prevent possible sexual transmission while at the games and after returning
 - Couples who want to try to get pregnant after going to the games should wait 8 weeks to 6 months
 depending on their situation.
- CDC and US Olympic Committee are working together to communicate risks and protective actions that people take before, during, and after travel to the games in Brazil.

LABORATORY TESTING

- Zika testing is recommended for
 - Anyone who has or recently experienced symptoms of Zika and lives in or recently traveled to an area with Zika.
 - o Anyone who has or recently experienced symptoms of Zika and had unprotected sex with a partner who lived in or traveled to an area with Zika.
 - o Pregnant women who live in or recently traveled to an area with Zika, with or without Zika symptoms.
- During the first two weeks after the start of illness, Zika virus disease can often be diagnosed by performing real-time reverse transcriptase polymerase chain reaction (rRT-PCR) on serum and urine.
 - o Zika virus rRT-PCR should be performed on serum and urine collected <14 days after onset of symptoms in patients with suspected Zika virus disease.
 - A positive Zika virus rRT-PCR confirms Zika virus infection. However, because Zika virus RNA in serum and urine decreases over time, a negative rRT-PCR does not rule out Zika virus infection; in this case, serologic testing should be performed.
 - o If Zika virus rRT-PCR results are negative for both specimens, serum should be tested by antibody detection methods.
- Serology assays can also be used to detect Zika virus-specific IgM and neutralizing antibodies, which typically develop toward the end of the first week of illness.
 - A positive IgM result does not always indicate Zika virus infection and can be difficult to interpret because cross-reactivity with related flaviviruses (e.g., dengue, Japanese encephalitis, West Nile, yellow fever) can occur.
 - A positive Zika virus IgM result may reflect previous vaccination against a flavivirus; previous infection with a related flavivirus; or current infection with a flavivirus, including Zika virus.
- Plaque-reduction neutralization testing (PRNT) can be performed to measure virus-specific neutralizing antibodies to confirm primary flavivirus infections and differentiate from other viral illnesses.
 - PRNT can be performed to measure virus-specific neutralizing antibodies to Zika virus, but neutralizing antibodies may still yield cross-reactive results in a person who was previously infected with another flavivirus, such as dengue, or has been vaccinated against yellow fever or Japanese encephalitis.
- Zika virus testing is performed at CDC, at some state and territorial health departments, and at some commercial laboratories. Healthcare providers should contact their state and local health department to facilitate testing. See the Testing for Zika Virus webpage for information on how to obtain Zika testing.

• Healthcare providers should work closely with the state or local health department to ensure that the appropriate test is ordered and interpreted correctly.

- o For specific testing recommendations, see <u>Testing for Pregnant Women</u> and <u>Testing for Infants and</u> Children.
- Laboratories processing clinical specimens for Zika virus diagnostic testing should, at a minimum, adhere to BSL2
 (biosafety level 2) precautions. All laboratories should perform a risk assessment to determine if there are certain
 procedures or specimens that may require higher levels of biocontainment. Suspicion that the specimen may
 contain a pathogen that requires BSL3 precautions (e.g., chikungunya virus), should be considered a significant risk
 factor.
- CDC is working to expand diagnostic testing capacity with both public and commercial partners in the United States.
- Each clinical scenario is unique, and healthcare providers should consider all available information when ordering a
 test for Zika virus infection including patient travel history, history of flavivirus infection, vaccination history,
 ultrasound findings, and the presence of symptoms. They should work with their state, local, and territorial health
 departments for assistance ordering laboratory tests and interpreting test results.

TYPES OF TESTS

- On February 26, 2016, the Food and Drug Administration (FDA) issued an Emergency Use Authorization (EUA) for a diagnostic tool for Zika virus that is being distributed to qualified laboratories and, in the United States, those that are certified to perform high-complexity tests.
 - The test, called the CDC Zika IgM Antibody Capture Enzyme-Linked Immunosorbent Assay (Zika MAC-ELISA), is intended to be used on serum and cerebrospinal fluid samples from people with a history of symptoms associated with Zika and/or people who meet the CDC Zika virus epidemiologic criteria (e.g., history of residence in or travel to a geographic region with active Zika virus transmission at the time of travel, or other epidemiologic criteria for which Zika virus testing may be indicated).
 - CDC has and continues to distribute the test to qualified laboratories in the Laboratory Response Network (LRN). The test is not available in US hospitals or primary care settings.
- On March 17, 2016, FDA issued an EUA for <u>a diagnostic tool</u> for the qualitative detection and differentiation of RNA from Zika virus, dengue virus, and chikungunya virus in human sera or cerebrospinal fluid, and for the qualitative detection of Zika virus RNA in urine and amniotic fluid.
 - The test, called the Trioplex Real-time RT-PCR Assay (Trioplex rRT-PCR), is intended to be used on specimens collected from people with a recent history of symptoms associated with Zika and/or people who meet the CDC Zika virus epidemiologic criteria (e.g., history of residence in or travel to a geographic region with active Zika virus transmission at the time of travel, or other epidemiologic criteria for which Zika virus testing may be indicated).
 - Testing is being performed by qualified laboratories designated by CDC and, in the United States, certified to perform high-complexity tests.
 - Because Trioplex rRT-PCR combines three tests (for Zika, dengue, and chikungunya) into one, it reduces
 costs and increases efficiency. Trioplex rRT-PCR has been adapted to be run on equipment that is common
 in public health labs in the United States and abroad.
- On April 28, 2016, FDA issued an EUA for a commercial assay for the qualitative detection of Zika virus RNA.
 - This test is the Focus Diagnostics, Inc., Zika Virus RNA Qualitative Real-time RT-PCR for use on acute serum only.
 - Unlike the Trioplex assay which tests for three viruses, the Focus Diagnostics RT-PCR is for the detection of Zika RNA in serum only.
- On May 13, 2016 the FDA issued an EUA for a commercial assay for the qualitative detection of RNA from Zika virus.

o This test is the Altona Diagnostics RealStar® Zika Virus RT-PCR Kit U.S. for the qualitative detection of RNA from Zika virus in serum or urine (collected alongside a patient-matched serum specimen).

- Unlike the Trioplex assay which tests for three viruses, the Altona Diagnostics RealStar® Zika Virus RT-PCR
 Kit is for the detection of Zika RNA in serum or urine.
- On June 17, 2016 the FDA issued an EUA for a commercial assay for the qualitative detection of RNA from Zika virus.
 - This test is Hologic, Inc. Aptima® Zika Virus assay for the qualitative detection of RNA from Zika virus in human serum and plasma specimens.
 - Unlike the Trioplex assay which tests for three viruses, the Aptima® Zika Virus assay is for the detection of Zika RNA in serum or plasma.

TESTING FOR PREGNANT WOMEN

- All pregnant women should be assessed for possible Zika virus exposure at each prenatal care visit. They should be asked if they:
 - o Traveled to or live in an area with active Zika virus transmission during their pregnancy or periconceptional period (the 6 weeks prior to last menstrual period or 8 weeks prior to conception).
 - Had sex without a condom with a partner who has traveled to or lives in an <u>area with active Zika virus</u> transmission. Condoms include male and female condoms.
- Possible exposure to Zika virus that warrants testing include:
 - Travel to or residence in an area with active Zika virus transmission, or
 - Sex (vaginal, anal, and oral sex or the sharing of sex toys) without a condom, with a person who traveled to, or lives in an area with Zika.
- Pregnant women who have a possible exposure to Zika virus are eligible for testing for Zika virus infection.
 - The type of testing recommended varies according to when a woman visits a provider relative to when her symptoms began or, if she is asymptomatic, the date of her last possible exposure to Zika virus.
- Asymptomatic pregnant women not living in an area with active Zika virus transmission whose potential exposure occurred less than 2 weeks prior to the current healthcare visit and women who had symptoms with an onset less than 2 weeks prior to the visit should receive Zika virus RT-PCR testing on both serum and urine.
 - o Positive rRT-PCR result confirms the diagnosis of recent maternal Zika virus infection.
 - o A negative rRT-PCR result does not rule out Zika virus infection because levels of Zika virus RNA in serum and urine decline over time.
 - For symptomatic pregnant women, Zika IgM and dengue IgM antibody testing should be performed immediately on the same specimen or a subsequently collected specimen.
 - Asymptomatic pregnant women should return 2 to 12 weeks after possible Zika virus exposure for Zika virus IgM antibody testing.
 - A negative result for both Zika IgM and dengue IgM indicate no recent Zika virus infection.
 - A positive or equivocal IgM result should be confirmed by PRNT.
- Symptomatic and asymptomatic pregnant women whose symptoms or possible exposure occurred 2 to 12 weeks prior to the healthcare visit should receive IgM testing.
 - Dengue IgM antibody testing is recommended only for symptomatic pregnant women.
 - o If the Zika virus IgM result is positive or equivocal, regardless of dengue IgM result, a reflex Zika virus RT-PCR should be performed on both serum and urine.
 - If the result is positive (on serum or urine), it confirms *recent zika virus infection*.
 - If the results are negative (on serum and urine) a PRNT should be performed.
 - A positive or equivocal dengue IgM result should also be confirmed by PRNT.

• Laboratory evidence of maternal Zika virus infection can include Zika virus RNA detected by rRT-PCR in any clinical specimen.

- Laboratory evidence of maternal Zika infection can include a positive or equivocal Zika virus IgM on serum or cerebrospinal fluid with a positive (≥10) PRNT titer for Zika virus together with a negative (<10) PRNT titer for dengue virus. Given that serology test results can be difficult to interpret, particularly in persons who were previously infected with or vaccinated against flaviviruses, and because the adverse outcomes caused by Zika virus infection during pregnancy are not fully described, pregnant women with laboratory evidence of recent flavivirus infection are considered to have possible Zika virus infection and should be monitored frequently
- Zika virus rRT-PCR can also be performed on amniotic fluid. Other testing that can be performed includes the following: 1) histopathologic examination and immunohistochemical staining of the placenta and umbilical cord, 2) Zika virus testing of frozen placental tissue and cord tissue, and 3) IgM and neutralizing antibody testing of infant serum and cord blood.

• For Pregnant Women with Ongoing Risk of Zika Virus Infection

- Ongoing risk for Zika virus exposure includes residence in or frequent travel to an area with active Zika virus transmission.
- Symptomatic pregnant women with ongoing risk of maternal Zika virus infection throughout their pregnancy should be tested for Zika virus infection.
- Asymptomatic pregnant women with ongoing risk of maternal Zika virus infection throughout their
 pregnancy should receive screening with serologic testing at the initiation of prenatal care and midsecond trimester. Local health officials should determine when to implement testing of asymptomatic
 pregnant women on the basis of information about local levels of Zika virus transmission and laboratory
 capacity.
- o If new symptoms consistent with Zika virus infection develop, a prior negative test for Zika virus does not rule out current infection. If new symptoms develop, a pregnant woman should be retested. During the first 14 days after symptom onset, viral RNA can often be identified in serum and urine, and RT-PCR is the preferred test to detect this. However, Zika virus RNA in urine and serum decreases over time, and a negative RT-PCR does **not** exclude Zika virus infection.

TESTING FOR INFANTS & CHILDREN

- Zika virus infection can be diagnosed by real-time reverse transcriptase polymerase chain reaction (rRT-PCR) for Zika virus RNA; OR Zika virus IgM and neutralizing antibodies.
 - It has not been established which test is most reliable for a diagnosis of congenital infection in newborns.
 Therefore, both should be performed.
 - A Zika virus rRT-PCR test should be performed on both infant serum and urine, and Zika virus immunoglobulin M (IgM) antibody enzyme-linked immunosorbent assay (ELISA) should be performed on infant serum.
 - Testing should be performed on infant specimens; testing of cord blood is no longer recommended.
 - Plaque-reduction neutralization testing (PRNT) needs to be performed in specimens that are positive for anti-Zika IgM antibodies to measure virus-specific neutralizing antibodies to differentiate Zika virus from infection with or vaccination for other flaviviruses.
 - If the infant's initial sample is IgM positive, but PRNT was not performed on the mother's sample, PRNT should be performed on the infant's initial sample. However, PRNT cannot distinguish between maternal or infant antibodies.
 - Maternal antibodies in the infant are expected to wane by 18 months.

 To confirm a congenital infection, PRNTs should be performed on a sample collected from an infant aged 18 months whose initial sample is IgM positive and neutralizing antibodies were detected by PRNT in either the infant's or mother's sample.

- o If cerebrospinal fluid (CSF) is obtained for other studies, rRT-PCR testing for Zika virus RNA and Zika virus IgM should be performed on CSF.
- Histopathologic evaluation of the placenta and umbilical cord, immunohistochemical staining on fixed tissue, and Zika virus rRT-PCR on fixed and frozen tissue can be performed.
- Zika virus infection can be confirmed in infants and children by performing reverse transcriptase-polymerase chain reaction (rRT-PCR) on serum within 7 days of symptoms onset.
 - Serologic assays can also be used to detect Zika virus-specific IgM and neutralizing antibodies soon after symptoms onset.
 - Evaluation of infants and children for acute Zika virus infection should include testing of serum and urine and may include cerebrospinal fluid (CSF) testing for Zika viral RNA, if samples were obtained as part of routine care. A CSF sample collected for the sole purpose of Zika rRT-PCR testing is not recommended.
- Laboratory evidence of Zika virus infection in an infant or child would include, in any clinical specimen, detectable Zika virus in culture, Zika virus RNA (by rRT-PCR) or antigen, or a clinical specimen positive for Zika virus IgM with confirmatory neutralizing antibody titers detected for Zika virus.
- Zika virus testing in newborns has several challenges.
 - o rRT-PCR tests may not detect Zika virus RNA in an infant or child who had Zika virus infection *in utero* if the period of viremia has passed.
 - Serologic tests for Zika virus can often be falsely positive because of general cross reactivity of IgM.
 - Plaque-reduction neutralization testing (PRNT) can be performed to measure virus-specific neutralizing antibodies to Zika virus, but maternal neutralizing antibodies (IgG) may cross the placenta. Because of this, PRNT testing cannot distinguish maternal from infant antibodies.
 - o It is important to work closely with state or territorial health departments to ensure the appropriate test is ordered and interpreted correctly.

CDC GUIDANCE & RECOMMENDATIONS

- CDC has developed guidance and recommendations on Zika for travelers, healthcare workers, and other groups. As new guidance and recommendations are developed and updated, they are posted on CDC's <u>Zika website</u>.
- CDC has interim guidelines for pregnant women and women of reproductive age with possible Zika virus exposure.
- Possible exposure to Zika virus that warrants testing include travel to or residence in an area with active Zika virus transmission, or sex (vaginal, anal, and oral sex, or the sharing of sex toys) with a partner who has traveled to or resides in an area with active Zika virus transmission without the use of a condom. Condoms include male and female condoms.
- Because there are limited data and experience with Zika virus in pregnancy, CDC continually evaluates any new or emerging data that may inform future recommendations. As more information becomes available, we will update the <u>CDC Zika website</u>.
 - o CDC has additional Q&As about Zika virus for healthcare providers.
- CDC has <u>interim guidelines</u> for healthcare providers caring for infants and children with possible Zika virus exposure. <u>Q&As</u> on these guidelines are also available.
- CDC has interim guidance for prevention of sexual transmission of Zika virus.
- CDC has <u>guidance</u> on response planning for Zika for district and school administrators in the continental United States and Hawaii.
- CDC has released <u>Health Alert Network</u> messages on Zika.

OBSTETRICAL HEALTHCARE PROVIDERS

CDC issued guidance and information to prevent Zika virus transmission and negative health outcomes, including
updated <u>interim guidance</u> for healthcare professionals for counseling patients about pregnancy planning and the
timing of pregnancy after possible exposure to Zika virus. <u>Updated interim guidance</u> was also issued for preventing
sexual transmission with information about how long men and women should consider using condoms or not
having sex after possible exposure or infection.

- For people who traveled to or live in an area with Zika and have a pregnant partner, CDC recommends
 using condoms every time during sex or not having sex during the pregnancy. To be effective, condoms
 must be used correctly, every time during sex. This includes vaginal, anal, and oral sex, and the sharing of
 sex toys.
- o For non-pregnant women and men who have been diagnosed with Zika virus or who have symptoms of Zika including fever, rash, joint pain or red eyes after possible exposure to Zika virus, CDC recommends
 - Women wait at least 8 weeks after their symptoms first appeared before trying to get pregnant.
 - Men wait at least 6 months after their symptoms first appeared to have unprotected sex.
- For non-pregnant women and men who traveled to an area with Zika but did not develop symptoms of Zika, consider using condoms or not having sex for at least 8 weeks after return in order to minimize risk of sexual transmission of Zika.
- Couples with men who live in an area with Zika but have not developed symptoms might consider using condoms or not having sex while there is active Zika transmission in the area.
- CDC has updated Interim Guidelines for Health Care Providers Caring for Pregnant Women with Possible Zika Virus Exposure.
 - Extended time frame for the laboratory test that detects presence of Zika virus in the blood of pregnant women with symptoms from <7 days to <14 days.
 - Added new recommendation to test for presence of Zika virus in the blood of some pregnant women without reported symptoms.
 - Added new recommendation to use the laboratory test that detects presence of Zika virus in the blood as
 a follow-up test after pregnant women test positive or have an equivocal test for an immune response to
 an infection that is likely to be Zika.
 - o The updated guidance also provides recommendations for female residents in areas with Zika virus.
- CDC released a <u>report</u> emphasizing the importance of healthcare personnel following practices, called Standard Precautions, to prevent the spread of infectious diseases such as Zika when caring for all patients, including pregnant patients in labor and delivery settings. Currently, there are no confirmed reports of Zika spreading from an infected patient to a health care provider or other patients. However, in labor and delivery units where health care personnel might come in contact with high volumes of body fluids, Standard Precautions to minimize contact with body fluids are important to reduce the possibility of spreading infectious diseases such as Zika.
- CDC continues to evaluate all available evidence and to update recommendations as new information becomes available. CDC's updated guidelines have been informed by our close collaboration with clinicians, professional organizations, state, tribal, local, and territorial health departments, and many other stakeholders.
- The American College of Obstetricians and Gynecologists and the Society for Maternal-Fetal Medicine (SMFM)
 have issued a <u>Practice Advisory</u> directed to obstetric providers about prevention strategies and clinical
 management of pregnant women.

ZIKA & PREGNANCY

• Counseling Pregnant Women

o Pregnant women should not travel to an area with Zika virus.

 Pregnant women who travel to or live in an area with active Zika virus transmission should strictly follow steps to avoid mosquito bites.

- Given the potential risks of maternal Zika virus infection, pregnant women who live in or travel to, or
 whose partner lives in or has traveled to an area with Zika should use a condom every time they have sex
 or not have sex during the pregnancy. To be effective, condoms must be used <u>correctly</u> (warning: this link
 contains sexually graphic images) from start to finish, every time you have sex. This includes vaginal, anal,
 and oral sex and the sharing of sex toys.
- o Recommendations will be updated as more information becomes available.

• Counseling Women Who are Considering Pregnancy

- For non-pregnant women and men who have been diagnosed with Zika virus or who have symptoms of Zika virus disease, including fever, rash, joint pain or red eyes, after possible exposure to Zika virus, CDC recommends
 - Women wait at least 8 weeks after their symptoms first appeared before trying to get pregnant.
 - Men wait at least 6 months after their symptoms first appeared to try to get their female partner pregnant.
 - People with Zika should use condoms for sex or not have sex during this time period, if they are
 concerned about spreading Zika virus to their sex partners. To be effective, condoms must be
 used every time, from start to finish during vaginal, anal, and oral sex and the sharing of sex toys.
- Non-pregnant women and men who traveled to an area with Zika but did not develop symptoms of Zika should consider using condoms or not having sex for at least 8 weeks after returning in order to minimize risk of sexual transmission of Zika virus.

• Counseling Women of Reproductive Age in Areas with Local Zika Virus Transmission

- A preconception counseling guide for healthcare providers counseling couples who are interested in conceiving and who live in an area with active Zika virus transmission is available <u>online</u>.
- Women of reproductive age with current or previous laboratory-confirmed Zika virus infection should be counseled that
 - CDC recommends waiting at least 8 weeks after their symptoms first appeared before trying to get pregnant.
 - There is no evidence that prior Zika virus infection poses an increased risk of birth defects in future pregnancies.
 - Zika virus usually remains in the blood of an infected person for about a week. There is no evidence that the virus will cause infection in a pregnancy that occurs after the virus is cleared from the mother's blood. The virus can be present in semen longer than in blood.
 - A woman thinking about pregnancy and recently recovered from Zika infection should talk to her doctor or other healthcare provider about planning her pregnancy.
- Healthcare providers should discuss reproductive life plans, including pregnancy intentions and timing with women of reproductive age in the context of the potential risks of Zika virus transmission.
 - A reproductive life plan helps a woman think about her goals for having or not having children
 and how to achieve these goals. A woman's plan depends on her personal goals. Reproductive
 life plan worksheets are available online.
- Healthcare providers should discuss strategies to prevent unintended pregnancy, including counseling on family planning and the correct and consistent use of effective contraceptive methods. Additionally, when choosing a contraceptive method, the prevention of sexually transmitted infections should also be considered, including the correct and consistent use of condoms.
- For women planning to become pregnant, healthcare providers should discuss the potential risk of Zika virus infection in pregnancy, the signs and symptoms associated with Zika virus disease, and when to seek

care if the patient develops symptoms of Zika virus disease. They should also emphasize strategies to prevent mosquito bites.

AMNIOCENTESIS

- Consideration of amniocentesis should be individualized for each patient's clinical circumstance. Healthcare providers should discuss the risks and benefits of amniocentesis with their patients.
- Similar to evaluation of other congenital infections, amniocentesis may be considered in the evaluation of potential Zika virus infection.
- It is unknown how sensitive or specific RT-PCR testing of amniotic fluid is for congenital Zika virus infection, whether a positive result is predictive of a subsequent fetal abnormality, and if it is predictive, what proportional of infants born after infection will have abnormalities. In addition, a negative result does not exclude congenital Zika virus infection.
- The optimal time to perform amniocentesis to diagnose congenital Zika virus infection is not known; Zika virus RNA has been detected in amniotic fluid as early as 4 weeks after maternal symptom onset, and as early as 17 weeks gestation.
- Amniocentesis performed ≥15 weeks of gestation is associated with lower rates of complications than those
 performed at earlier gestational ages (≤14 weeks of gestation).
- The exact timing of amniocentesis should be individualized based on the patient's clinical circumstances. Referral to a maternal-fetal medicine or infectious disease specialist with expertise in pregnancy management may be warranted. Risks and benefits of performing the amniocentesis should be discussed with the patient.

PRENATAL DIAGNOSIS OF MICROCEPHALY

- Microcephaly and intracranial abnormalities have been demonstrated in pregnancies with known Zika virus
 infection. Hence, additional ultrasounds might provide an opportunity to identify findings consistent with fetal Zika
 virus infection and offer pregnant women the option of amniocentesis to test for Zika virus RNA.
- Microcephaly can be diagnosed during pregnancy with ultrasound. Microcephaly is most easily diagnosed by ultrasound in the late second trimester or early third trimester of pregnancy.
- Fetal ultrasound is generally performed in pregnancies between 18-20 weeks of gestation to assess fetal anatomy as part of routine obstetrical care.
- Although microcephaly and intracranial calcifications are typically detected during ultrasounds in the late second
 and early third trimester of pregnancy, these findings might be detected as early as 18-20 weeks gestation.
 However, detection by prenatal ultrasound can be challenging at this gestational age due to fetal position and fetal
 motion artifact.
- The optimal time to perform ultrasound screening for fetal microcephaly is not known. In the absence of
 microcephaly, the presence of intracranial calcifications before 22 weeks gestation might suggest a risk for the
 future development of microcephaly.
- Zika virus infection is a cause of microcephaly and other severe fetal brain defects. Brain abnormalities reported in
 infants with laboratory-confirmed congenital Zika infection include microcephaly and disrupted brain growth.
 Some infants with possible Zika virus infection have been found to have intracranial calcifications and
 abnormalities of the eye.
 - In one published report of two infants with Zika virus RNA detected by RT-PCR, brain anomalies detected
 on ultrasound included corpus callosal and vermian dysgenesis, enlarged cisterna magna, severe
 unilateral ventriculomegaly, agenesis of the thalami, cataracts, intracranial and intraocular calcifications.
- Ultrasound is performed during pregnancy when medical information is needed. It has been used during pregnancy for many years and has not been associated with adverse maternal, fetal, or neonatal outcomes.

Ultrasound operators are trained to use the lowest power for the minimum duration of time to obtain the
needed information. There is consensus among various national and international medical organizations
(American College of Radiology, American College of Obstetricians and Gynecologists, and the Society for
Maternal Fetal Medicine) that ultrasound is safe for the fetus when used appropriately.

- The accuracy of ultrasound to detect microcephaly in the setting of maternal Zika virus is not known and will depend on many factors, such as the timing of maternal infection relative to the timing of screening, severity of microcephaly, patient factors (e.g., obesity), gestational age, the equipment used, and the expertise of the person performing the ultrasound.
 - Because the absence of fetal microcephaly and intracranial calcifications on ultrasound at one point in pregnancy does not exclude future microcephaly, additional ultrasounds may be considered at the discretion of the healthcare provider. As we get more information specifically related to Zika virus infection and microcephaly, we expect that more specific guidance for women and their healthcare providers will be developed.
- The sensitivity of prenatal ultrasound for detection of microcephaly depends on a range of factors (e.g., timing of
 ultrasound screening, severity of microcephaly, patient factors). In a study of fetal microcephaly not caused by Zika
 virus infection, prenatally ultrasound-diagnosed microcephaly correlated with neonatal microcephaly
 approximately 57% of the time.
- Fetal MRI is not a screening tool and should be used only to answer specific questions raised by ultrasound or used
 in occasional specific high-risk situations. Interpretation of fetal MRI requires specialized expertise and has limited
 availability and accessibility in the United States.

PEDIATRIC HEALTHCARE PROVIDERS

- CDC has interim guidelines for healthcare providers in the United States caring for infants and children with possible congenital or perinatal Zika virus infection.
 - Update: Interim Guidance for the Evaluation and Management of Infants with Possible Congenital Zika
 Virus Infection United States, August 2016
- These guidelines include recommendations for the evaluation, testing, and management of infants with possible congenital Zika virus infection. These interim guidelines will be updated as more information becomes available.
- The baseline prevalence of congenital microcephaly is difficult to determine because of underreporting, and the
 inconsistency of clinical criteria used to define microcephaly. Although population-based estimates of congenital
 microcephaly in Brazil vary, the number of infants with microcephaly currently being reported in Brazil is greater
 than would be expected.
- Although Zika virus RNA has been detected in breast milk, transmission of Zika infection through breastfeeding has
 not been documented. Based on available evidence, the benefits of breastfeeding infants outweigh any theoretical
 risk. CDC encourages mothers with Zika virus infection and mothers living in areas with Zika to breastfeed their
 infants.
- We do not know if a newborn who gets Zika virus at birth will develop microcephaly after birth. Babies can develop microcephaly after birth if their head growth slows or fails to develop after birth. There have been no reports of Zika virus infection around the time of birth leading to microcephaly in infants.
- The prognosis for infants with congenital Zika virus infection is not known.

ZIKA VIRUS TRANSMISSION IN INFANTS & CHILDREN

- Zika virus can be transmitted from a pregnant woman to her fetus during pregnancy or around the time of birth. We do not know how often Zika transmission occurs during pregnancy or around the time of birth.
- Congenital or intrauterine transmission of Zika virus occurs when a woman is infected with Zika virus during her
 pregnancy, but before delivery, and the virus passes to the fetus.

• Perinatal transmission of Zika virus occurs when a woman is infected with the Zika virus within 2 weeks of delivery, and the virus passes to the infant at or around the time of delivery.

- When an infant acquires Zika virus disease perinatally, the infant may develop symptoms such as maculopapular rash, conjunctivitis, arthralgia (joint symptoms), and fever.
- Zika virus RNA has been identified in breast milk. No evidence of Zika virus infection associated with breastfeeding has been reported.
 - Current evidence suggests that the benefits of breastfeeding outweigh the theoretical risks of Zika virus infection transmission through breast milk. CDC encourages mothers with Zika virus infection and mothers living in areas with Zika virus to breastfeed their infants.

BIRTH DEFECTS

- In a published review of existing evidence, CDC scientists concluded that Zika virus infection during pregnancy can cause microcephaly and other severe fetal brain defects. Scientists continue to study the full range of other potential health problems that Zika virus infection during pregnancy may cause.
- No treatment is currently available for Zika virus infection. Care for these infants is focused on diagnosing and
 managing conditions that are present, monitoring the child's development over time, and addressing problems as
 they arise.
- From what we know about severe <u>microcephaly</u>, a range of neurologic sequelae has been reported (e.g., intellectual disability, hearing loss, vision loss, and seizures). These problems can range from mild to severe, are often life-long, and in some cases can be life-threatening.
- Microcephaly is diagnosed when an infant's head is smaller than expected as compared to infants of the same age
 (or gestational age) and sex. Although a universally accepted definition of microcephaly does not exist,
 microcephaly is most often defined as head circumference (occipitofrontal circumference) greater than 2 standard
 deviations below the mean, or less than the 3rd percentile based on standard growth charts.
 - o For infants diagnosed with microcephaly, head size correlates with underlying brain size. However, these measurements do not consistently predict long-term sequelae.
 - Neurologic sequelae may include seizures, vision or hearing problems, and developmental disabilities.
 Sequelae vary with the extent of brain disruption.
 - Causes of congenital microcephaly may include genetic conditions such as chromosomal abnormalities or maternal exposures (e.g., alcohol, mercury, or radiation) during pregnancy. Other maternal infections that have been associated with microcephaly include cytomegalovirus (CMV), herpes simplex virus, rubella virus, lymphocytic choriomeningitis virus (LCMV), *Treponema pallidum* (i.e., syphilis), and *Toxoplasma* gondii.
- Head circumference (HC) and occipitofrontal circumference (OFC) are the same. These terms can be used interchangeably.
 - Head circumference measurements should be taken using a tape measure that cannot be stretched. The tape is securely wrapped around the widest possible circumference of the head, 1-2 finger widths above the eyebrow on the forehead and at the most prominent part of the back of the head. It may be helpful to have the parent or nurse hold the infant's arms. Ideally, the measurement should be taken 3 times and the largest measurement to the nearest 0.1 cm should be used.
 - Although head circumference measurements may be influenced by molding and other factors related to
 delivery, the measurements should be taken on the first day of life because commonly-used birth head
 circumference reference charts by age and sex are based on measurements taken before 24 hours of age.
 The most important factor is that the head circumference is carefully measured and documented. If
 measurement within the first 24 hours of life is not done, the head circumference should be measured as
 soon as possible after birth.

POTENTIAL OUTCOMES & PROGNOSIS

• There is limited information on neurocognitive outcomes in neonates if they are exposed to Zika virus during labor and delivery or after birth.

- o Perinatal transmission of Zika virus infection has been reported. However information is limited to two cases: one of these infants was asymptomatic and the other had thrombocytopenia and a diffuse rash.
- Evidence from other flaviviruses, such as West Nile virus and dengue virus, indicate that transmission has
 resulted in findings in the neonate ranging from no symptoms to severe illness (including fever,
 thrombocytopenia, and hemorrhage).
- The spectrum of clinical features that might be observed in infants who acquire Zika virus during the perinatal period is currently unknown.
- The prognosis for infants with congenital Zika virus infection is not known.
- For infants with congenital Zika virus infection, care is focused on diagnosing and managing conditions that are present, monitoring the child's development over time, and addressing problems as they arise.
- Information on long-term outcomes among infants and children with acute Zika virus disease is limited. Thus, until
 more evidence is available to inform recommendations, routine pediatric care is advised for these infants and
 children.
- Most children infected with Zika virus are asymptomatic or have mild illness, similar to the findings seen in adults with Zika virus infection.
 - o Treatment is supportive; this includes rest and fluids to prevent dehydration.
 - Non-steroidal anti-inflammatory drugs (NSAIDS) should not be used until dengue is ruled out as a cause of illness and should be avoided in children aged < 6 months.
 - Aspirin is not recommended for use with acute viral illnesses due to the risk of Reye's syndrome.
- In general, the risk for Guillain-Barré syndrome from any cause appears to increase with increasing age. GBS has been reported following Zika virus infection, although a causal link has not been established.
 - It is unclear how often GBS following Zika virus infection has occurred in children; one report from Brazil
 refers to 6 patients, aged 2–57 years, with neurologic syndromes (4 with GBS and 2 with acute
 disseminated encephalomyelitis) after laboratory-confirmed Zika virus infection; no further data are
 available.
 - Deaths due to Zika virus infection appear to be very rare at all ages.

CLINICAL GUIDANCE

• Congenital Zika Virus Infection

- CDC updated its interim guidance for the evaluation and testing of infants with possible congenital Zika virus infection (published on February 26, 2016). In the <u>new guidance</u>, CDC also provides recommendations for the outpatient management of infants with laboratory evidence of possible Zika virus infection, with or without apparent associated birth defects.
- o In the updated guidance, CDC includes recommendations for the care of infants with possible congenital Zika virus infection throughout the first 12 months of life.
- All infants born to mothers with laboratory evidence of Zika virus infection during pregnancy should receive a comprehensive physical exam, head ultrasound to assess the brain's structure, standard newborn hearing assessment, and lab testing for Zika virus.
- Within one month of age, infants with laboratory evidence of Zika virus infection should also receive a detailed eye exam and a more in-depth hearing exam.
- Additional recommendations for follow-up and services for infants born to women with evidence of Zika virus infection during pregnancy depend on whether these infants have birth defects consistent with congenital Zika syndrome.
- Although data on outcomes associated with congenital Zika virus infection are limited, experiences with other congenital infections can provide insight to guide clinical management until more data emerge.

Infants with congenital infections, such as congenital cytomegalovirus and congenital rubella syndrome, can develop a range of disabilities later in life, including hearing loss, seizures, and neurodevelopmental delays, even without signs of infection at birth.

- Families of affected infants will require support and referrals for information and services. There is likely to be a disproportionate burden on families with limited access to medical care and barriers to services.
- Testing is recommended for infants born to mothers with laboratory evidence of possible Zika virus infection. Testing is also recommended for infants with signs of congenital Zika syndrome at birth, such as brain abnormalities, if the mother has an epidemiologic link (lived in or traveled to an area with Zika and/or had sex without a barrier method to prevent infection with a partner who lived in or traveled to an area with Zika).
 - Testing should be performed within the first 2 days of life, if possible. Later testing will make it difficult to determine if perinatal or postnatal infection has occurred.
 - Because the types of services needed to care for infants with congenital Zika syndrome are complex, CDC recommends coordinated care through a multidisciplinary team and established medical home.
 - As a critical component of patient care and early identification of any delays, families should be empowered to be active participants in their child's monitoring and care.
 Resources for families can be found at CDC's website.
- For infants born to mothers with risk factors for maternal Zika virus infection (travel to or residence in an
 area of active Zika virus transmission or sex without a condom with a partner with travel to or residence
 in such an area) for whom maternal testing was not performed before delivery, assessment of the infant,
 including comprehensive physical exam and careful measurement of head circumference should be
 performed.
 - Maternal diagnostic testing should be performed and testing of the placenta for Zika virus PCR should be considered.
 - If an infant appears clinically well, further evaluation and infant testing can be deferred until maternal test results are available. However, if there is concern about infant follow-up, infant testing should be performed before hospital discharge.
- o In many cases, infant testing results will not be available before hospital discharge. If test results are not available before hospital discharge, infants should be presumed to have congenital Zika virus infection until test results are available. Infants with confirmed and probable Zika virus infection should be managed in the same way, according to guidance.
- If an infant has confirmed or probable congenital Zika virus infection, a thorough physical examination should be performed, including careful measurement of the head circumference, length, weight, assessment of gestational age, neurologic assessment, postnatal head ultrasound, and standard newborn hearing screen.
 - Infants with confirmed or probable congenital Zika virus infection should have a comprehensive ophthalmologic exam and hearing assessment by auditory brainstem response (ABR) testing before 1 month of age.
- Infants with laboratory evidence of Zika virus infection but without abnormalities consistent with congenital Zika syndrome should receive routine care including monitoring of head (occipitofrontal) circumference, length, and weight.
 - Additionally, developmental monitoring should occur at every visit, and age-appropriate standardized validated developmental screening should occur at 9 months.
 - A vision screening, including visual regard, should be performed at each well-child visit, and referral to an ophthalmologist should be made for any caregiver or provider concerns.
 - Infants who passed an initial ABR should be referred for behavioral audiological diagnostic testing at age 9 months, or sooner for any hearing concerns.
- If an infant has confirmed or probable congenital Zika virus infection, or test results and findings
 consistent with congenital Zika syndrome, consultations are recommended with a pediatric neurologist, a
 pediatric infectious disease specialist, an ophthalmologist, an endocrinologist, and a clinical geneticist or
 dysmorphologist.

A complete blood count, platelet count, and liver function tests should also be conducted.
 Testing for other infectious or genetic causes for abnormalities is also recommended. Advanced neuroimaging should be considered in consultation with a neurologist.

- For infants with negative Zika test results and without suspected abnormalities, healthcare providers should continue with routine pediatric care including measurement of growth and development, and appropriate evaluation and follow-up for any clinical findings that arise.
 - If the newborn has abnormal findings consistent with congenital Zika syndrome but negative testing, diagnostic testing for other causes of the newborn's conditions should be performed including testing for other infectious or genetic causes for abnormalities.
- If a mother had Zika virus infection during pregnancy but the newborn tests negative for Zika virus:
 - If the newborn does not have abnormal findings on examination, the infant should receive routine pediatric care including measurement of growth and development, and appropriate evaluation and follow-up for any clinical findings that arise.
 - If the newborn has abnormal findings on examination, diagnostic testing for other causes of the newborn's conditions should be performed including testing for other congenital viral infections if indicated.

• Pediatric Infection

- o Acute Zika virus disease should be suspected in an infant or child aged <18 years who
 - 1) traveled to or resided in an area with Zika virus within the past 2 weeks and
 - 2) has ≥1 of the following manifestations: fever, rash, conjunctivitis, or arthralgia.
- Because transmission of Zika virus from mother to infant during delivery is possible, acute Zika virus disease should also be suspected in an infant during the first 2 weeks of life
 - 1) whose mother traveled to or resided in an affected area within 2 weeks of delivery and
 - 2) who has ≥1 of the following manifestations: fever, rash, conjunctivitis, or arthralgia.
- Arthralgia can be difficult to detect in infants and young children and can manifest as irritability, walking with a limp (for ambulatory children), difficulty moving or refusing to move an extremity, pain on palpation, or pain with active or passive movement of the affected joint.

WHAT CDC IS DOING

- CDC's <u>Emergency Operations Center</u> (EOC) was activated January 22, 2016, and moved to a level 1 activation—the highest level on February 8, 2016. The EOC is the command center for monitoring and coordinating the emergency response to Zika, bringing together CDC scientists with expertise in arboviruses like Zika, reproductive health, birth defects, developmental disabilities, and travel health. Their work includes:
 - Developing laboratory tests to diagnose Zika.
 - Conducting studies to learn more about learn more about Zika and its effects during pregnancy and the potential link between Zika and Guillain-Barré syndrome.
 - Conducting a study to evaluate persistence of Zika virus in semen and urine among male residents of the United States.
 - Publishing and disseminating a new report on the persistence of Zika virus in blood among five pregnant women reported to the US Zika Pregnancy Registry.
 - Publishing and disseminating a new report estimating contraceptive use among sexually active US women of reproductive age where mosquito-borne Zika virus transmission is possible.
 - Monitoring and reporting cases of Zika, which will help improve our understanding of how and where Zika
 is spreading.
 - o Providing guidance to travelers and Americans living in areas with current outbreaks.
 - Providing on-the-ground support in American Samoa, Brazil, Colombia, Guam, the Marshall Islands,
 Panama, Puerto Rico, Trinidad and Tobago, and the US Virgin Islands.
- CDC's EOC is currently home to hundreds of CDC staff working in collaboration with local, national, and international response partners to analyze, validate, and efficiently exchange information about the outbreak.

• The EOC has resources to rapidly transport diagnostic kits, clinical specimens that will be tested for Zika virus, and personnel.

- The EOC is serving as CDC's command center for monitoring and coordinating the emergency response to Zika, including sending CDC staff and the procurement and management of all equipment and supplies that CDC responders may need during deployment.
- CDC is sending staff to assist with the response senior leaders, vector control, emergency management, logistician, epidemiology/surveillance, data entry, pregnancy and birth defects, blood safety specialists, etc.

DOMESTIC ACTIVITIES

- CDC is working with public health partners and with state health departments to
 - o Alert healthcare providers and the public about Zika.
 - o Post travel guidance.
 - Provide state health laboratories with diagnostic tests.
 - o Monitor and report cases of Zika.
 - Publish and disseminate guidelines to inform testing and treatment of people with suspected or confirmed 7ika
 - o Publish and disseminate conclusions on the causal association between Zika and microcephaly.
- In response to local transmission of Zika transmission in Florida, at Florida's request, CDC is sending a CDC Emergency Response Team (CERT) with experts in Zika virus, pregnancy and birth defects, vector control, laboratory science, and risk communication to assist in the response.
 - CDC has updated its <u>Interim Zika Response Plan (CONUS and HI)</u> and has released <u>the Zika Community</u>
 <u>Action Response Toolkit (Z-CART)</u> to help states with risk communication and community engagement
 when local transmission is identified.
- On April 1, 2016, CDC hosted the Zika Action Plan Summit for state and local health officials. The Summit aimed to
 - o Provide officials with information and tools to improve Zika preparedness and response within their states and jurisdictions.
 - o Increase knowledge on the latest Zika science, including implications for pregnant women.
 - o Increase knowledge of best communication practices, including crisis and risk communication principles.
 - Accelerate readiness for local Zika transmission through training and technical assistance to states to help establish and support surveillance and share best practices for vector control.
 - o Identify possible gaps in preparedness and response at the federal, state, and local levels, and to help begin to address possible gaps.
- CDC released guidance documents to assist in preparedness and response planning for state, local, and territorial public health officials.
 - o <u>Zika Communication Planning Guide for States</u>
 - o Interim CDC Recommendations for Zika Vector Control in the Continental United States
- CDC's health security plans are designed to effectively monitor for disease, equip diagnostic laboratories, and support mosquito control programs both in the United States and around the world.
- CDC is creating and distributing Zika Prevention Kits for affected US territories (Puerto Rico, US Virgin Islands, and American Samoa).

ACTIVITIES IN PUERTO RICO

- CDC continues to deploy staff to Puerto Rico to support all aspects of the Zika outbreak including:
 - o Evaluating vector control interventions and implementing an island-wide vector control program
 - Improving diagnostic tests

 Supporting active surveillance of pregnant women infected with Zika virus in Puerto Rico, and follow-up efforts for infants born to these women for up to 3 years

- Establishing the first surveillance system for cases of Guillain-Barré syndrome, as well as an investigation to better determine the association between Zika virus infection and the neurologic condition
- Conducting community engagement activities with the purpose of and implementing mosquito control programs at the local level
- Educating pregnant women about Zika virus infection prevention and distributing Zika prevention kits
- Conducting research on Zika virus, such as the persistence of the virus in different bodily fluids
- Evaluating four Zika prevention interventions (mass media, community outreach, Zika prevention kit, and vector control) among pregnant women recruited from Women, Infants, and Children (WIC) program across Puerto Rico
- CDC supports the development of health education campaigns for Puerto Rico:
 - "This is How We Stop Zika" or "Deten el Zika", a multi-media communication campaign that provides steps for pregnant women and communities to follow to protect themselves from Zika virus infection, mainly by taking actions to prevent mosquito bites and avoiding potential sexual transmission of the virus. The campaign includes 2 websites (English www.helpstopzika.org and Spanish www.detenelzika.org), a 30 sec- TV PSA, radio PSAs, billboards, posters, newspaper ads, community engagement toolkit and social media content
 - Through the domestic readiness campaign, CDC has placed ads (some using PRDH logos, and some Gates campaign logos) focusing on the four protective behaviors people can take on billboards, in shopping malls, bar and restaurant restrooms, on buses, bus/metro stops, movie theaters, newspapers, and panel displays in grocery stores to reach pregnant women and their families
- The CDC Foundation collaborates with CDC and multiple organizations to provide a range of contraceptive options to women and their partners who want to delay or avoid pregnancy during the Zika outbreak in Puerto Rico.

INTERNATIONAL ACTIVITIES

- CDC is working in laboratories and in dozens of countries, with ministries of health, and with partners around the
 world to develop a deeper understanding of Zika virus. We are also helping to prevent, control, and respond to the
 Zika outbreak, along with outbreaks of other diseases like chikungunya, dengue fever, malaria, yellow fever, and
 other vector-borne diseases. CDC is working through our country offices, our programs, and with international
 partners to
 - o Alert healthcare providers and the public about Zika.
 - o Provide health laboratories with diagnostic tests. Through the CDC's Emergency Operations Center, CDC is assisting countries with Zika testing by supplying them with reagents for molecular diagnostic lab testing.
 - Monitor and report cases of Zika, which will help improve our understanding of how and where Zika is spreading.
 - Learn more about Zika and its effects during pregnancy as well as potential link between Zika and Guillain-Barré syndrome.
- CDC is committed to global health security. We help build the capacity of even the most vulnerable countries to detect, prevent, and respond to public health emergencies within their own borders.
- CDC staff are providing essential lab assistance, including
 - Organizing and triaging requests for Zika virus PCR reagents, needed for Zika diagnostics testing, from CDC's 10 Global Disease Detection Regional Centers and around the world.
 - Standing up and operating regional Zika virus laboratory diagnostic capabilities at two of the most critical Global Disease Detection Regional Centers - the Central America Regional Center in Guatemala and the SE Asia Regional Center in Thailand
 - Developing a next-generation diagnostics card for acute febrile illness that tests samples of up to 8 people for 30 pathogens simultaneously, including Zika, delivering results in less than 3 hours
- Through its 24/7 Global Disease Detection Operations Center, regional Global Disease Detection Centers, Country Offices, and global Field Epidemiology and Laboratory Training Programs, CDC is working with governments,

ministries of health, and international partners to conduct rigorous surveillance for new and emerging infections, identify and characterize new pathogens, develop and evaluate new laboratory methods, and train disease detectives in the countries in which they operate.

- CDC's Global Disease Detection (GDD) Operations Center operates 24/7 and is continually carrying out event-based surveillance to monitor this outbreak globally.
 - This center, in collaboration with Zika subject matter experts and international partners and governments, has been conducting event-based surveillance to monitor spread of Zika from Brazil to other areas in the Americas since May 2015, sharing this information to coordinate the response.
 - The GDD program, launched in 2004, was one of the first ways CDC systematically began helping countries build the systems they need to prevent, detect and respond to health threats. Its regional centers are currently working with governments and international partners to provide data from the field back to CDC's Global Disease Detection Operations Center for global surveillance.
 - Because of this work we are able to know:
 - Where Zika is spreading in the Americas
 - Where (in what regions) we are seeing increased numbers of microcephalic babies
 - Where there are significant upticks in Guillain-Barré syndrome.
 - O CDC's GDD Center, part of the Central America Region office located in Guatemala City has been critical in:
 - Ensuring that emergency operations centers in countries with Zika in Central America and beyond are equipped and ready to activate and perform, and that different government agencies in each country know how to collaborate across programs and agencies on the response.
 - Helping Colombia, El Salvador, Guatemala, Honduras, and Panama with lab testing for Zika and chikungunya.
 - Instituting four functions vital to disease detection and surveillance in the countries it serves:
 - Developing a common testing platform and protocol for influenza;
 - Drafting and testing of national emergency action plans;
 - Training field epidemiologists and lab technicians through the Field Epidemiology and Lab Training Programs (FETP and FELTP)
 - Strengthening laboratory capabilities by:
 - Supporting Zika virus testing in Guatemala, laboratory diagnostic training in South and Central America, and specimen transport in the Latin American region; and sharing protocols and procedures with labs in Lima, Peru to strengthen overall ability to accurately and quickly test for selected diseases.
 - Supporting countries with Zika in vector surveillance, control strategies, and insecticide resistance testing in coordination with the Pan American Health Organization (PAHO)
- CDC is collaborating with Colombia's Instituto Nacional de Salud (INS) on a project called Proyecto Vigilancia de Embarazadas con Zika (VEZ), which involves close follow-up of pregnant women and their infants in sites with the highest number of Zika-infected pregnant women. Through this collaboration, CDC and INS hope to better understand the full range of potential health problems that congenital Zika virus infection may cause, the risk of poor outcomes among fetuses/infants of women infected with Zika virus during pregnancy, and the time of pregnancy during which Zika virus infection poses the highest risk to the fetus.
- In an effort to better understand Zika and its effects during pregnancy, the Brazilian Ministry of Health (MOH) and PAHO are investigating.
 - o PAHO invited CDC to provide technical assistance to the Brazil MOH for its investigation of microcephaly and the possible association with Zika virus infection by collaborating on studies.
 - CDC regularly communicates with representatives from PAHO and the Brazil MOH to discuss the investigation and laboratory testing options.

• CDC has offered to test samples from the microcephaly cases for serologic evidence of Zika virus infection until in-country capacity can be established.

- CDC's FETP and FELTP programs provide real-time training to capable "disease detectives" and laboratory specialists in these countries who can identify and target disease.
- CDC's Central America Field Epidemiology Training Program (CA FETP) includes national field epidemiology training programs in Belize, Costa Rica, Dominican Republic, El Salvador, Haiti, Guatemala, Honduras, and Panama.
 - These countries are coordinated through REDCEC (Red Centroamericana de Epidemiologia de Campo;
 Central American Network of Field Epidemiology), under the umbrella organization of the Council of
 Health Ministers of Central America and the Dominican Republic (COMISCA).
 - o Brazil, Mexico, Colombia, and additional South American countries have independent FETPs.

CDC FOUNDATION

- At CDC's request, the CDC Foundation activated two funds on February 10, 2016, to help with the Zika response: the U.S. Emergency Response Fund and the Global Disaster Response Fund.
- These funds allow CDC to better prepare for and respond to crisis situations, such as Zika, by providing flexibility to meet needs that would not otherwise be met through federally appropriated funds.
- Financial support raised through these funds will enhance CDC's response in a number of ways, such as extend the agency's ability to alert healthcare providers and the public about Zika; fund Zika prevention kits with educational materials for pregnant women in Puerto Rico and other high-risk areas; protect travelers with guidance and information; support state health laboratories with diagnostic tests; and detect and report cases to help prevent further spread.
- The CDC Foundation recently announced a partnership with CDC and multiple donors to create Zika prevention kits for pregnant women in the US territories of the Commonwealth of Puerto Rico, the US Virgin Islands (USVI), and American Samoa. The purpose of these Zika prevention kits is to inform pregnant women about Zika, its risks, and how to avoid infection, while providing an initial supply of prevention tools. The initial donations included mosquito repellent, mosquito dunks, and condoms.
- Walgreens also <u>recently announced</u> a collaboration with CDC and the CDC Foundation on Zika education and prevention in Puerto Rico. In addition, Walgreens made a \$100,000 donation to the CDC Foundation to aid its Zika education and prevention efforts.
- On April 1, 2016, the CDC Foundation, along with Bayer and SC Johnson, provided donor support for the Zika Action Summit at CDC, which was attended by federal and state government representatives as well as representatives from associations, philanthropies, and the private sector.
- Educating communities and empowering women on how to prevent Zika virus transmission is the focus of a new collaborative effort by CDC, the Pan American Health Organization (PAHO) and the CDC Foundation, aimed mainly at pregnant women in U.S. territories and the Americas. The Bill & Melinda Gates Foundation is supporting these efforts, which include a comprehensive health campaign on Zika prevention and surveys on risk perception and knowledge gaps in the Americas, as well as community engagement on mosquito control especially to protect pregnant women from Zika. These initiatives will be funded by a \$1.5 million grant to the CDC Foundation and PAHO.
- The CDC Foundation also announced collaborations with CDC and multiple organizations, including <u>Bayer</u>, <u>Allergan</u>, <u>Medicines360</u>, <u>Upstream USA</u>, and <u>Merck</u>, to provide a range of contraceptive options to women and their partners who want to delay or avoid pregnancy during the Zika outbreak in Puerto Rico.
- CDC Foundation board members Dikembe Mutombo and Gary Cohen <u>authored an op-ed</u> that appeared on CNBC about the need for private sector funding in the Zika response.
- The CDC Foundation recently <u>announced</u> the launch of the Zika prevention communication campaign in Puerto Rico. The campaign, titled "This is How We Stop Zika," provides steps for pregnant women and communities to

follow to protect themselves from Zika virus infection, mainly by taking actions to prevent mosquito bites and avoiding potential sexual transmission of the virus. However, as mosquito season is expected to continue into December, the CDC Foundation urgently needs \$1.5 million in funding to continue this vitally important campaign in Puerto Rico.

- CDC Foundation President & CEO Dr. Judith Monroe authored <u>an op-ed</u> on CNBC that emphasizes three actions our country must undertake to beat back Zika.
- On August 25, the CDC Foundation announced that the Zika Contraception Access Network (Z-CAN) is now operational. The Network is providing women in Puerto Rico with a full range of contraceptive options free of charge on the same day of their healthcare service. Z-CAN was established by the CDC Foundation to address an urgent need to improve contraception access in Puerto Rico during the Zika outbreak. The program gives women who want to delay or avoid pregnancy an effective means to do so, and the option to prevent the devastating, lifelong consequences of severe birth defects Zika virus can cause.
 - To date, the Z-CAN team has trained approximately 160 physicians and 260 ancillary staff in Puerto Rico to counsel and provide a full range of contraception to women wanting to delay or avoid pregnancy during the Zika outbreak. In addition, the CDC Foundation team has secured contraceptive product donations, established a supply chain for distribution of contraceptive products across the island, as well as created a system to reimburse physicians.
 - Donations of funding and contraceptive product commitments have been crucial to reduce program
 implementation costs, but additional funding is required to fully execute this effort at the speed and scale
 required for the Zika response. The CDC Foundation currently has funding to reach approximately 14,000
 women in Puerto Rico. With a goal of \$18 million in funding, which is the estimated cost of caring for two
 infants with microcephaly, this effort could be expanded to serve tens of thousands of more women
 during the time of Zika.